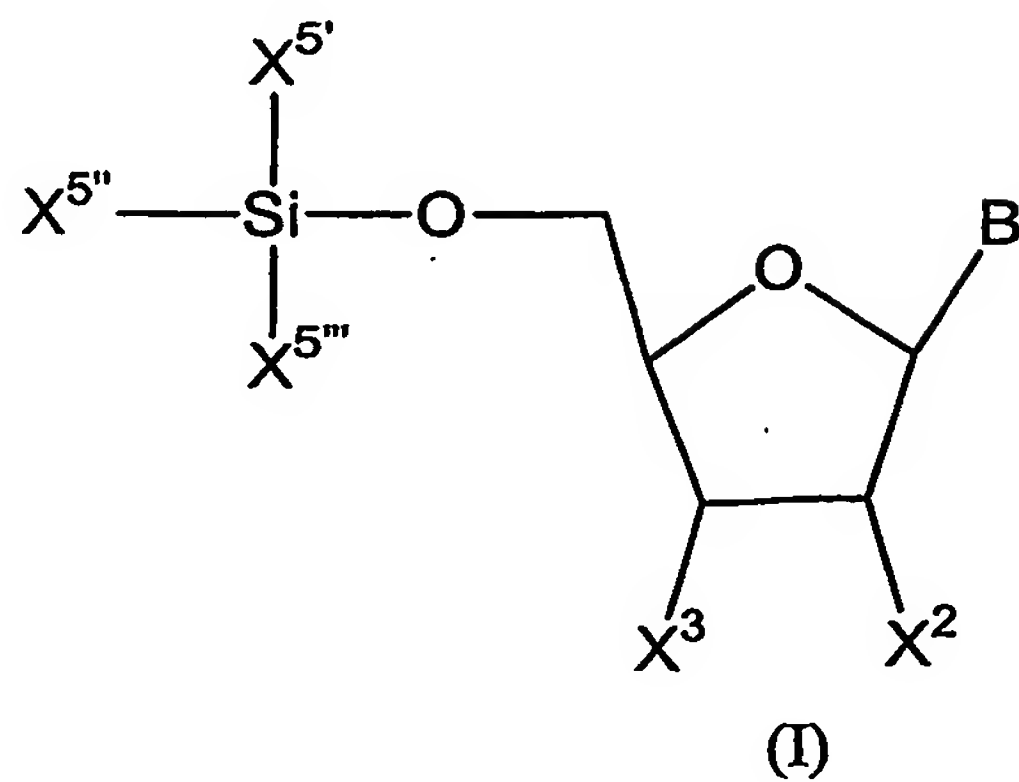
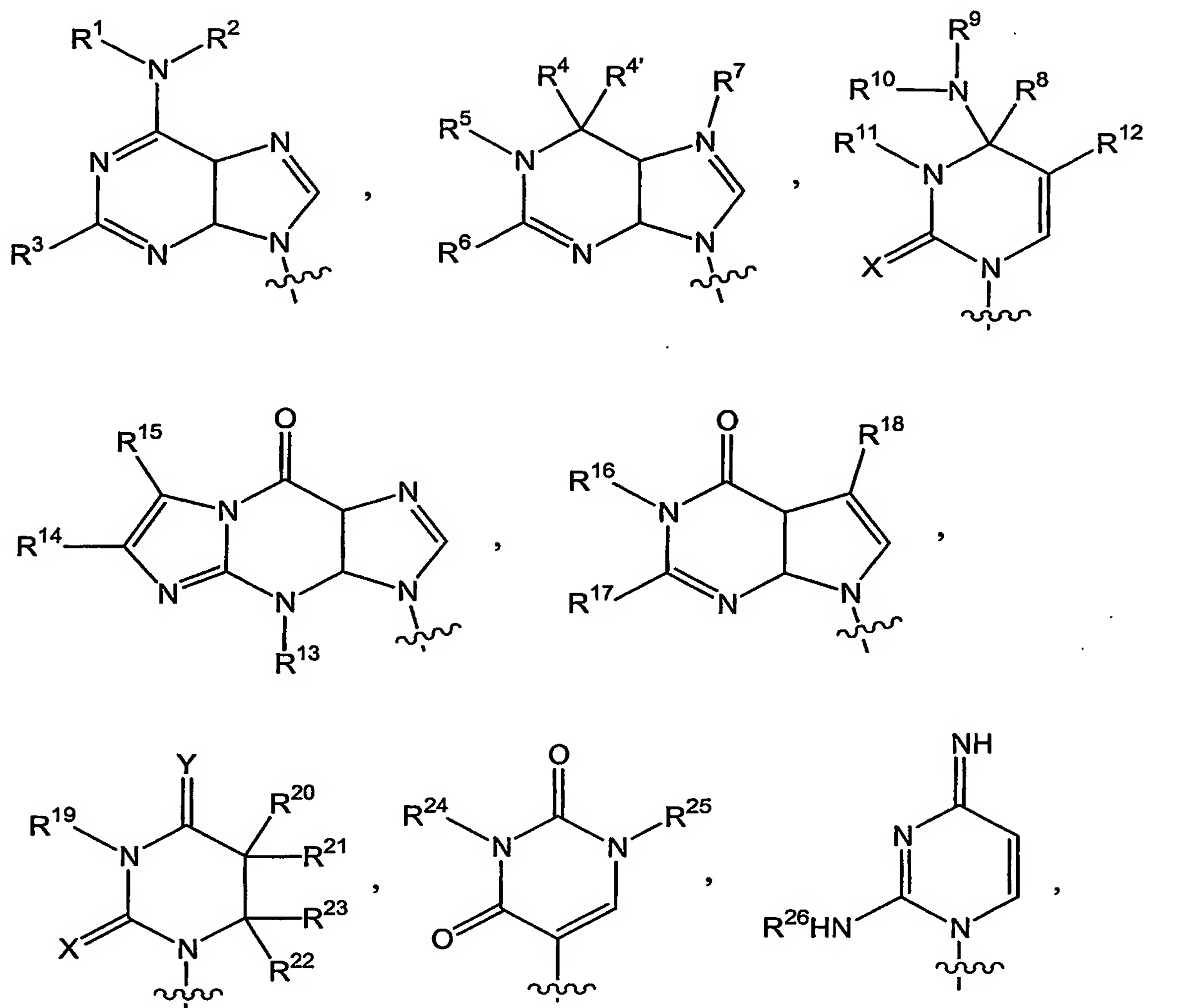


WHAT IS CLAIMED IS:

1. A protected monomer having a formula (I)



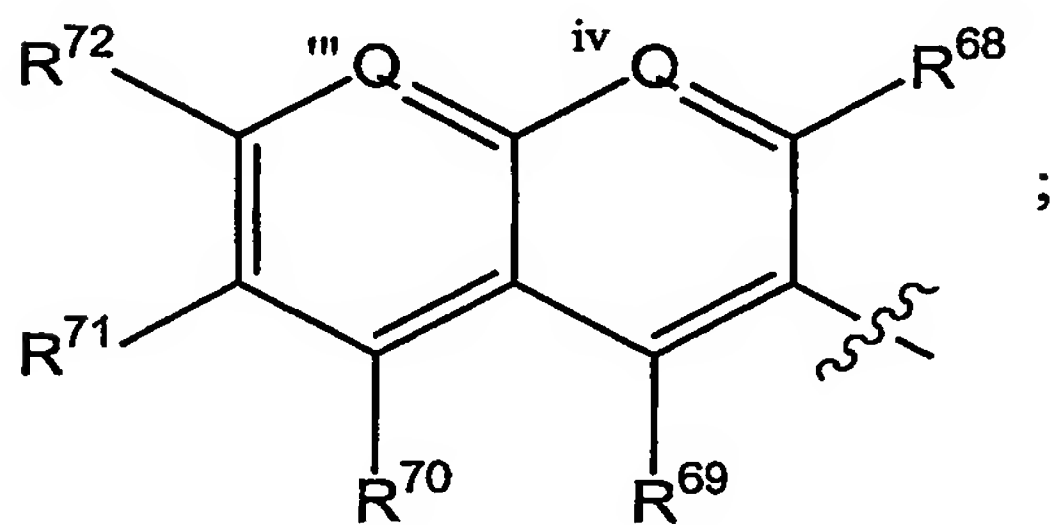
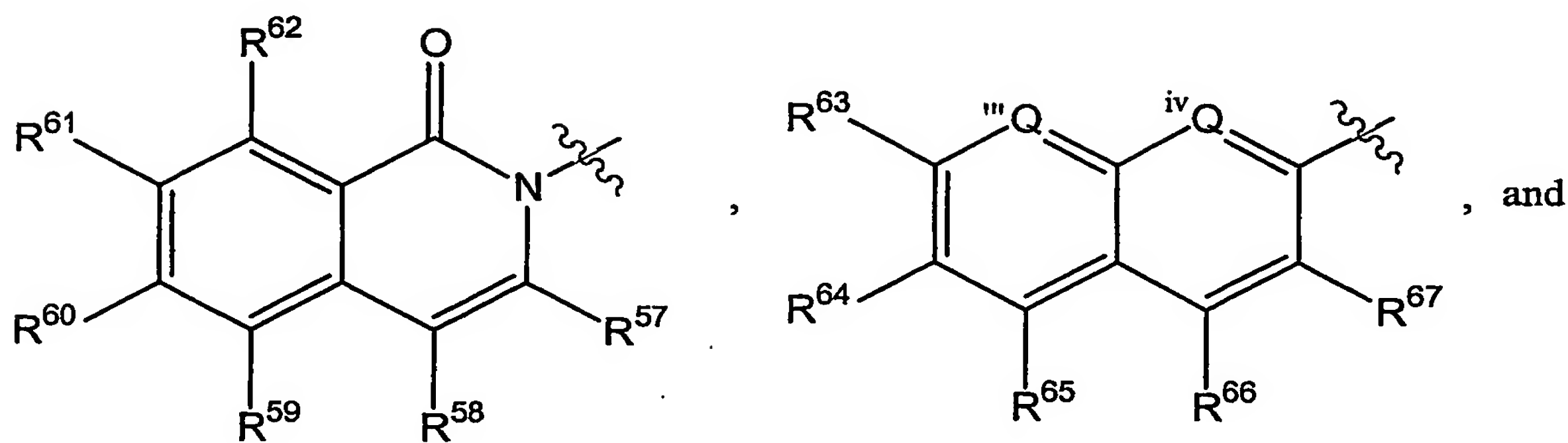
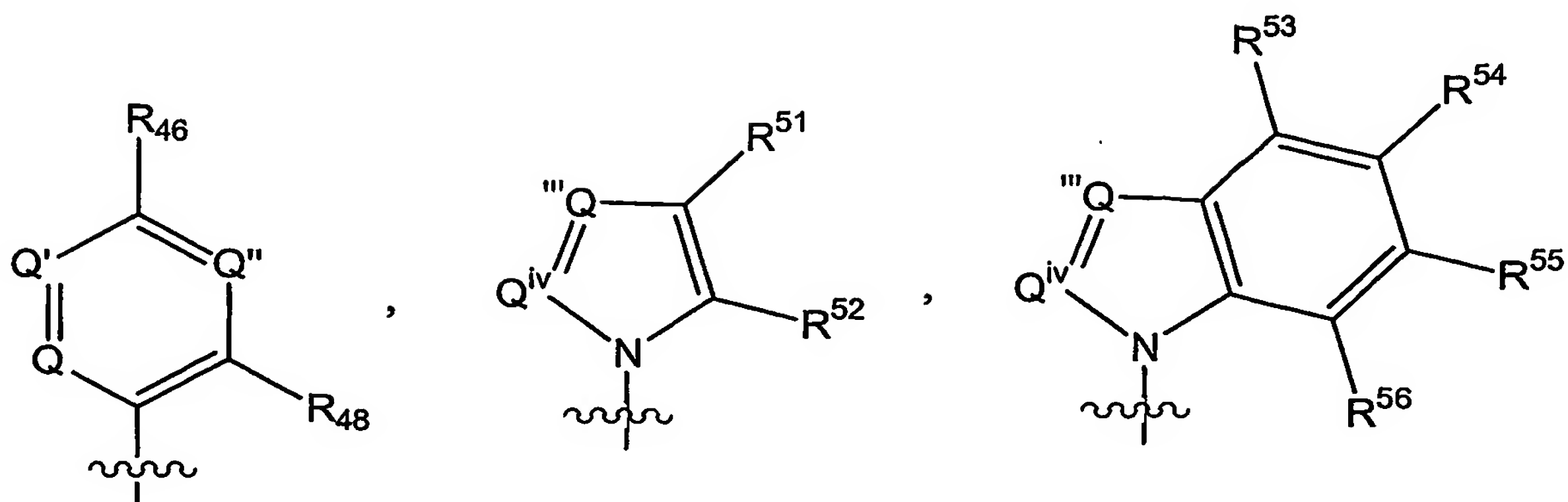
- 5 wherein,
 B is selected from the group consisting of:



anthracenyl, pyrenyl,

5

10



X^2 is an ortho ester protecting group, hydrogen, ethers, alkyl ethers, esters, halogens, protected amines, or protected hydroxyl moieties;

5 X^3 is $-O-P(OR^{27})N(R^{28})_2$ or $-O-L-R^{29}$;

$X^{5'}$, $X^{5''}$, $X^{5'''}$ include at least one alkoxy or siloxy substituent;

R^1 is hydrogen or C_1 - C_4 alkyl;

R^2 is hydrogen, C_1 - C_4 alkyl, or C_2 - C_6 alkenyl optionally substituted with hydroxy, or $C(O)NHR^a$;

10 R^3 is hydrogen, halo, C_1 - C_4 alkyl, C_1 - C_4 thioalkoxy, NH_2 , NHR^b , or NR^bR^c ;

R^4 when taken together with $R^{4'}$ forms oxo, or R^4 when taken together with R^5 forms a double bond between the carbon and nitrogen atoms to which they are attached;

$R^{4'}$ when taken together with R^4 forms oxo, or is O^- ;

R^5 is hydrogen, C_1 - C_4 alkyl, or when taken together with R^4 forms a double bond
5 between the carbon and nitrogen atoms to which they are attached;

R^6 is hydrogen, halo, NH_2 , NHR^b , or NR^bR^c ;

R^7 is an unshared electron pair, or C_1 - C_4 alkyl;

R^8 when taken together with R^9 forms a double bond between the carbon and nitrogen
atoms to which they are attached, or R^8 when taken together with R^{11} forms a double bond
10 between the carbon and nitrogen atoms to which they are attached;

R^9 is hydrogen, C_1 - C_4 alkyl, or when taken together with R^8 forms a double bond
between the carbon and nitrogen atoms to which they are attached;

R^{10} is hydrogen or is absent;

R^{11} is hydrogen, C_1 - C_4 alkyl, or when taken together with R^8 forms a double bond
15 between the carbon and nitrogen atoms to which they are attached;

R^{12} is hydrogen, formyl, or C_1 - C_4 alkyl optionally substituted with hydroxy or protected
hydroxy;

R^{13} and R^{14} are each independently hydrogen or C_1 - C_4 alkyl;

R^{15} is hydrogen, C_1 - C_4 alkyl, or $(CH_2)_nCH(R^d)CH(NHR^e)(COOR^g)$;

20 R^{16} is hydrogen or C_1 - C_4 alkyl;

R^{17} is halo, NH_2 , NHR^b , or NR^bR^c ;

R^{18} is cyano, $C(=NH)NH_2$, or $CH_2NH(R^h)$;

R^{19} is hydrogen, or C_1 - C_4 alkyl;

R^{20} is:

25 (i) hydrogen;

(ii) hydroxy or protected hydroxy;

(iii) C_1 - C_4 alkoxy optionally substituted with $COOR^f$; or

(iv) C_1 - C_4 alkyl optionally substituted with hydroxy and/or $COOR^f$, NH_2 , NHR^m , or
 $CONH_2$;

30 R^{21} is hydrogen, or when taken together with R^{23} forms a double bond between the
carbon atoms to which they are attached;

R^{22} is hydrogen;

R^{23} is hydrogen, or when taken together with R^{21} forms a double bond between the
carbon atoms to which they are attached;

R^{24} and R^{25} are each, independently, hydrogen or C_1 - C_4 alkyl;

R^{26} is $(CH_2)_nCH(R^d)CH(NHR^e)(COOR^f)$;

R^{27} is C_1 - C_6 alkyl optionally substituted with cyano, or C_2 - C_6 alkenyl;

R^{28} is C_1 - C_{10} alkyl;

5 R^{29} is a liquid or solid phase support reagent;

Q is N or CR^{44} ;

Q' is N or CR^{45} ;

Q'' is N or CR^{47} ;

Q''' is N or CR^{49} ;

10 Q^{iv} is N or CR^{50} ;

R^{44} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, a ligand, a tethered ligand, or when taken together with R^{45} forms $-OCH_2O-$;

15 R^{45} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, a ligand, a tethered ligand, or when taken together with R^{44} or R^{46} forms $-OCH_2O-$;

R^{46} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, a ligand, a tethered ligand, or when taken together with R^{45} or R^{47} forms $-OCH_2O-$;

20 R^{47} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, a ligand, a tethered ligand, or when taken together with R^{46} or R^{48} forms $-OCH_2O-$;

25 R^{48} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, a ligand, a tethered ligand, or when taken together with R^{47} forms $-OCH_2O-$;

R^{49} , R^{50} , R^{51} , R^{52} , R^{53} , R^{54} , R^{57} , R^{58} , R^{59} , R^{60} , R^{61} , R^{62} , R^{63} , R^{64} , R^{65} , R^{66} , R^{67} , R^{68} , R^{69} , R^{70} , R^{71} , and R^{72} are each independently selected from hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_2 - C_6 alkynyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, $NC(O)R^{17}$, or $NC(O)R^0$;

30 R^{55} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_2 - C_6 alkynyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, $NC(O)R^{17}$, or $NC(O)R^0$, or when taken together with R^{56} forms a fused aromatic ring which may be optionally substituted;

R^{56} is hydrogen, halo, hydroxy, nitro, protected hydroxy, NH_2 , NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_2 - C_6 alkynyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl, $NC(O)R^{17}$, or $NC(O)R^o$, or when taken together with R^{55} forms a fused aromatic ring which may be optionally substituted;

5 X is O, S, or Se;

Y is O or S;

L is $-C(O)(CH_2)_qC(O)-$, or $-C(O)(CH_2)_qS-$;

10 Provided that R^1 , R^2 , and R^3 cannot all be hydrogen; further provided that when R^5 is hydrogen, R^6 cannot be NH_2 , $NH(\text{protecting group})$, or $NH(iBu)$; further provided that when R^{12} is hydrogen and R^8 and R^{11} together form a double bond between the carbon and nitrogen atoms to which they are attached, R^9 and R^{10} cannot both be hydrogen; further provided that when X and Y are O, R^{19} is hydrogen, and R^{21} and R^{23} together form a double bond between the carbon atoms to which they are attached, R^{20} cannot be hydrogen or CH_3 ;

15 R^a is glycyl, threonyl, or norvalyl, each of which may optionally be partially or fully protected;

R^b is C_1 - C_6 alkyl or a nitrogen protecting group;

R^c is C_1 - C_6 alkyl;

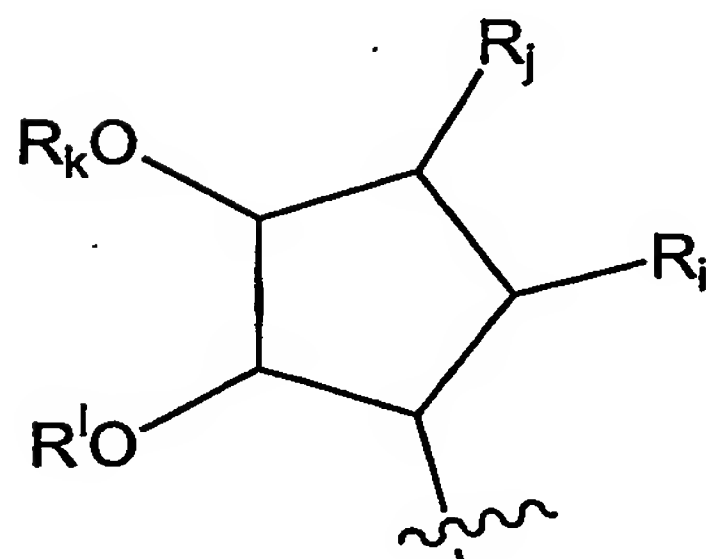
R^d is hydrogen, hydroxy, protected hydroxy, or OOH ;

R^e is hydrogen, a nitrogen protecting group, or $COOR^g$;

20 R^f is hydrogen, or C_1 - C_6 alkyl;

R^g is C_1 - C_{10} alkyl;

R^h is hydrogen, or



25 R^i and R_j when taken together forms a double bond between the carbon atoms to which they are attached, or R^i and R_j when taken together form $-O-$ between the carbon atoms to which they are attached;

R^k and R^l are each, independently, hydrogen, a hydroxyl protecting group, a sugar, or a fully or partially protected sugar;

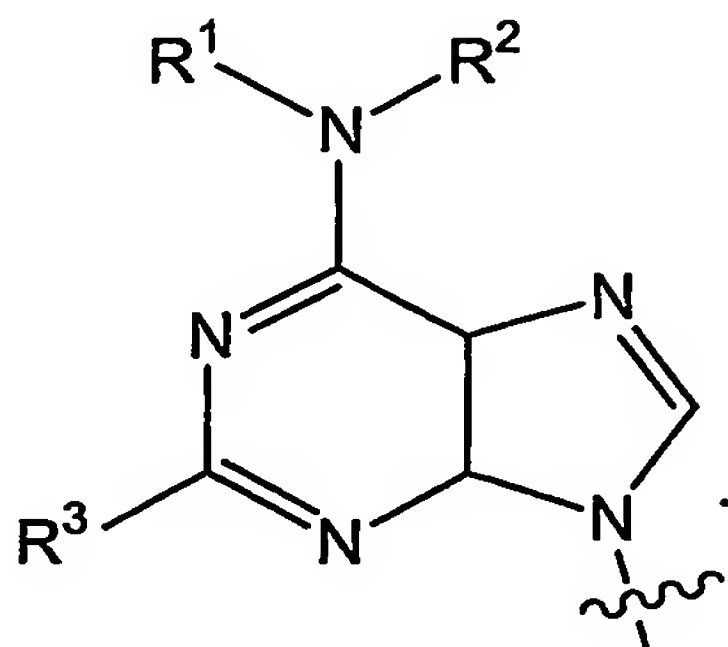
R^m is C_1 - C_4 alkyl optionally substituted with $COOH$;

R^o is alkyl optionally substituted with halo, hydroxy, nitro, protected hydroxy, NH_2 ,
5 NHR^b , or NR^bR^c , C_1 - C_6 alkyl, C_2 - C_6 alkynyl, C_6 - C_{10} aryl, C_6 - C_{10} heteroaryl, C_3 - C_8 heterocyclyl,
 $NC(O)R^{17}$, or $NC(O)R^o$;

n is 1-4; and

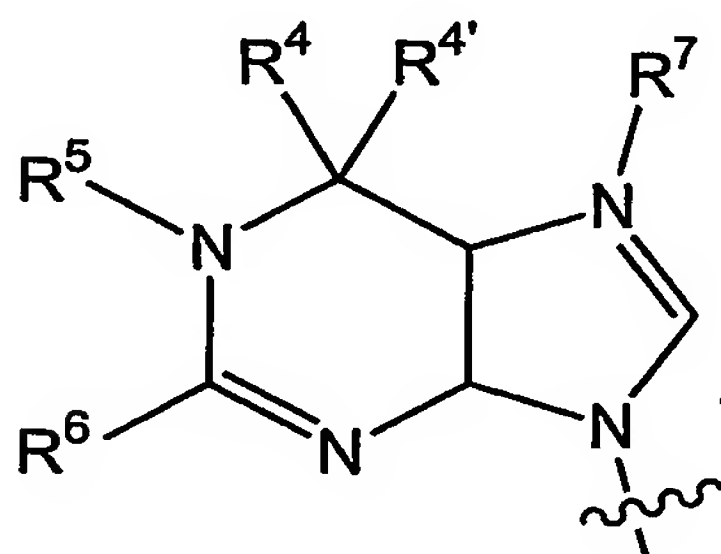
q is 0-4.

2. The monomer of claim 1, wherein B is:

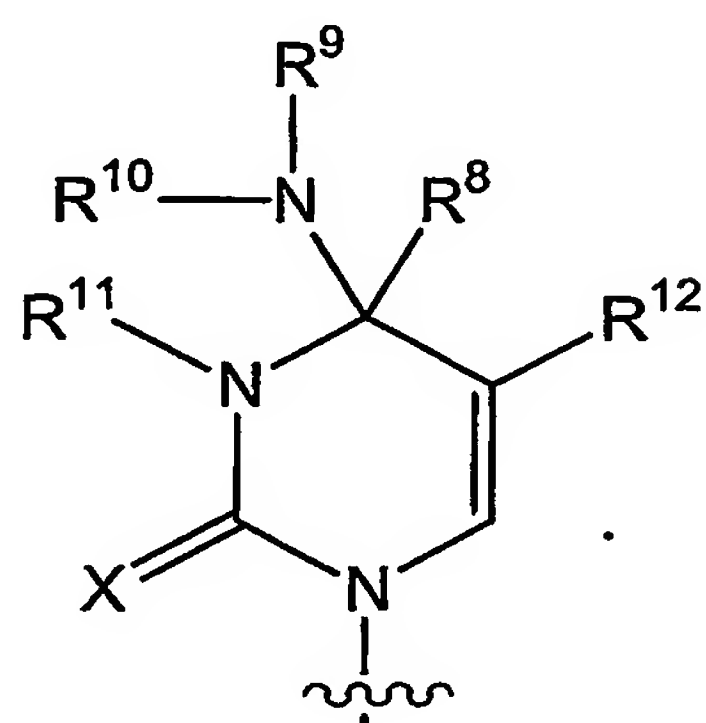


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3. The monomer of claim 1, wherein B is:

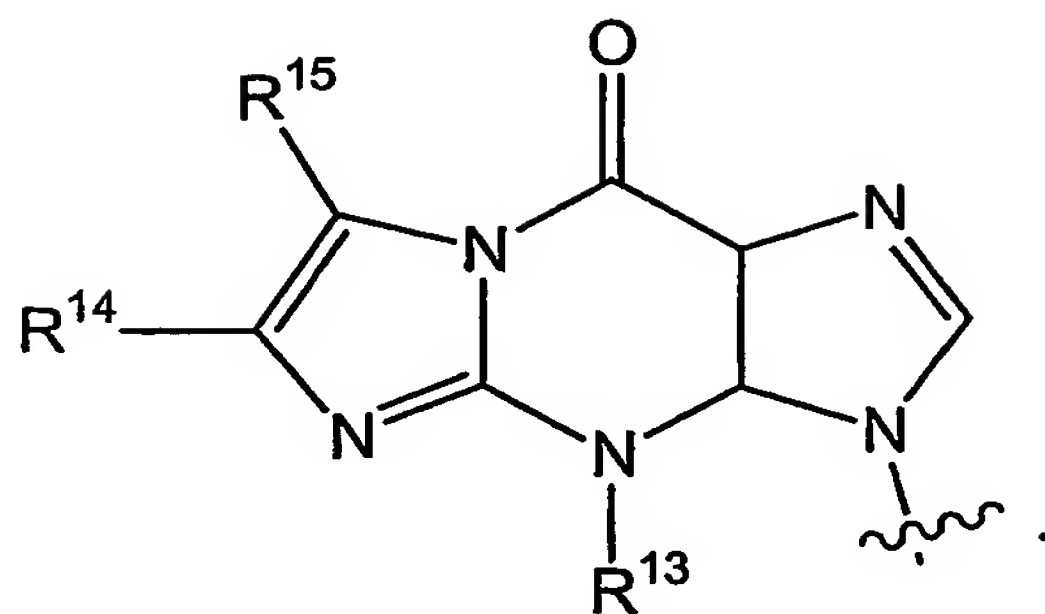


4. The monomer of claim 1, wherein B is:



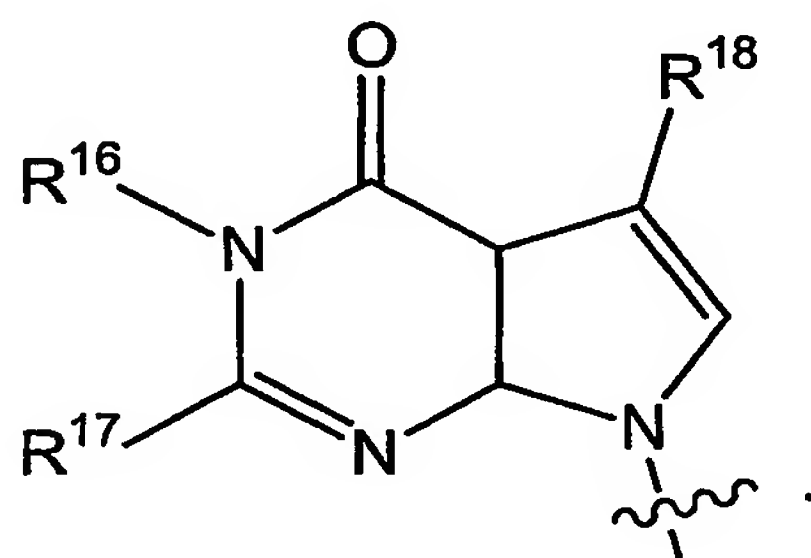
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5. The monomer of claim 1, wherein B is:

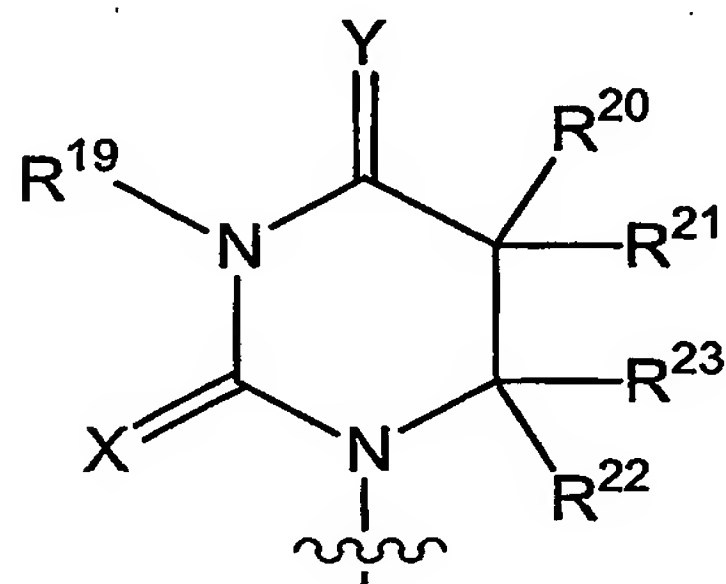


6. The monomer of claim 1, wherein B is:

5

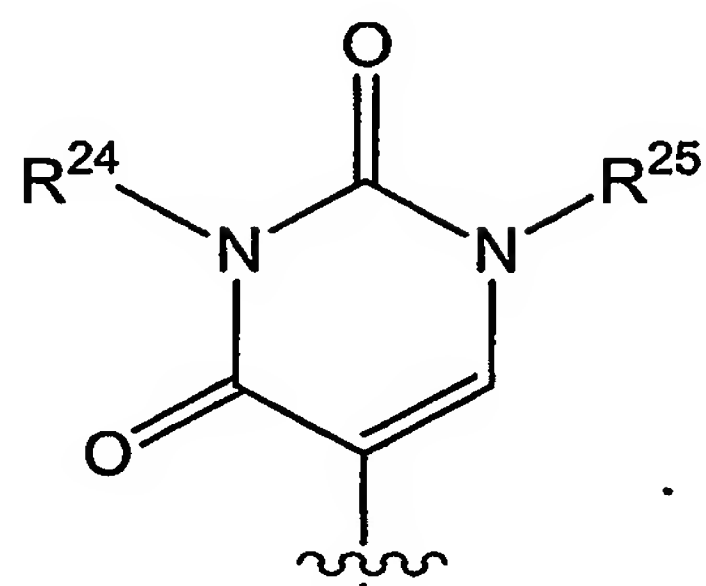


7. The monomer of claim 1, wherein B is:



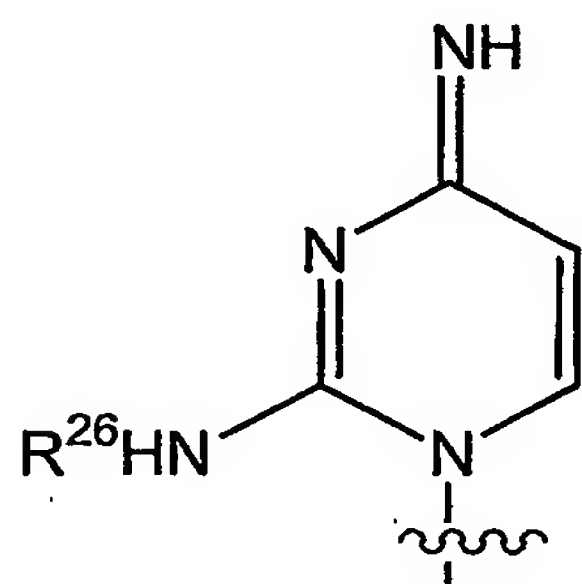
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8. The monomer of claim 1, wherein B is:

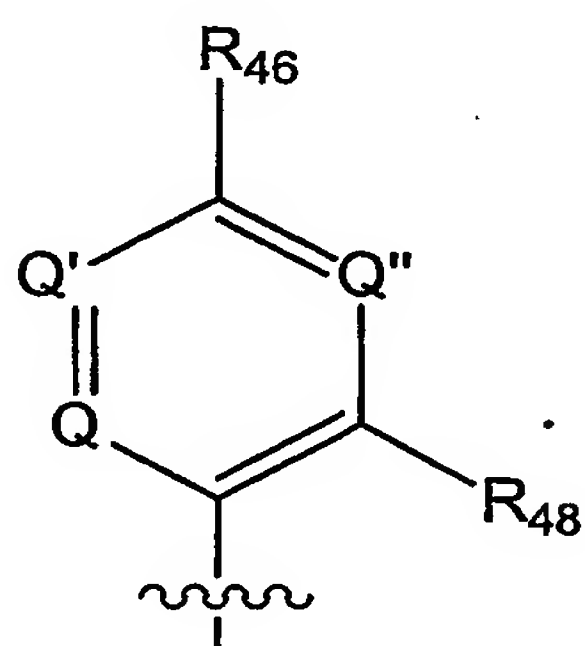


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9. The monomer of claim 1, wherein B is:

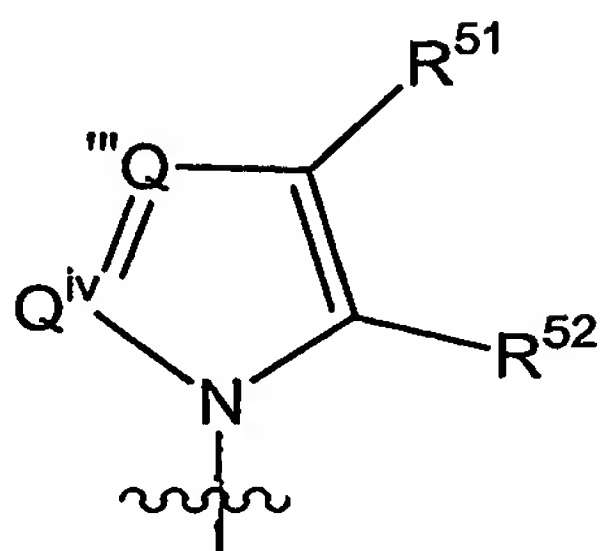


10. The monomer of claim 1, wherein B is:



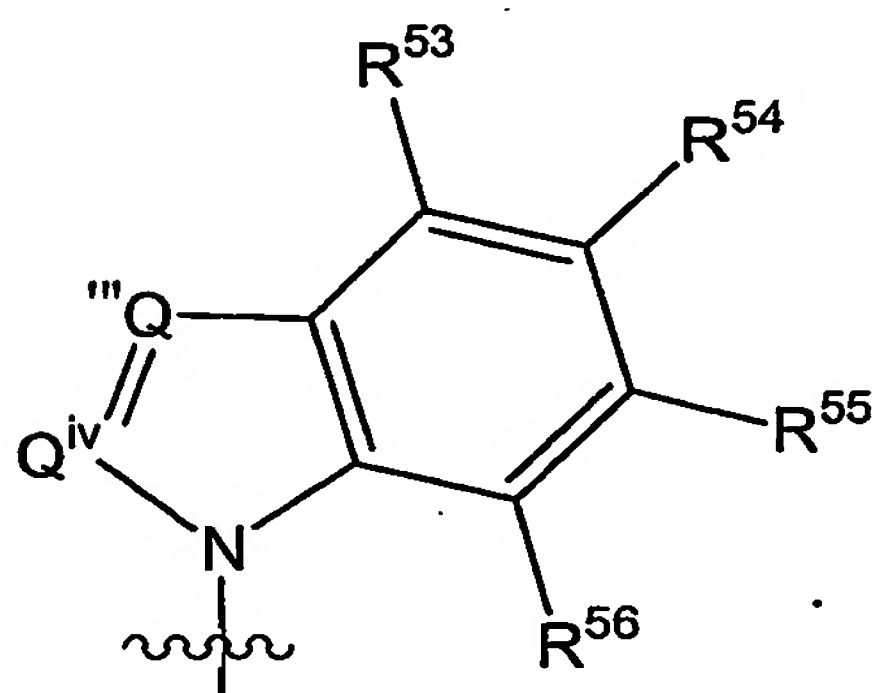
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11. The monomer of claim 1, wherein B is:

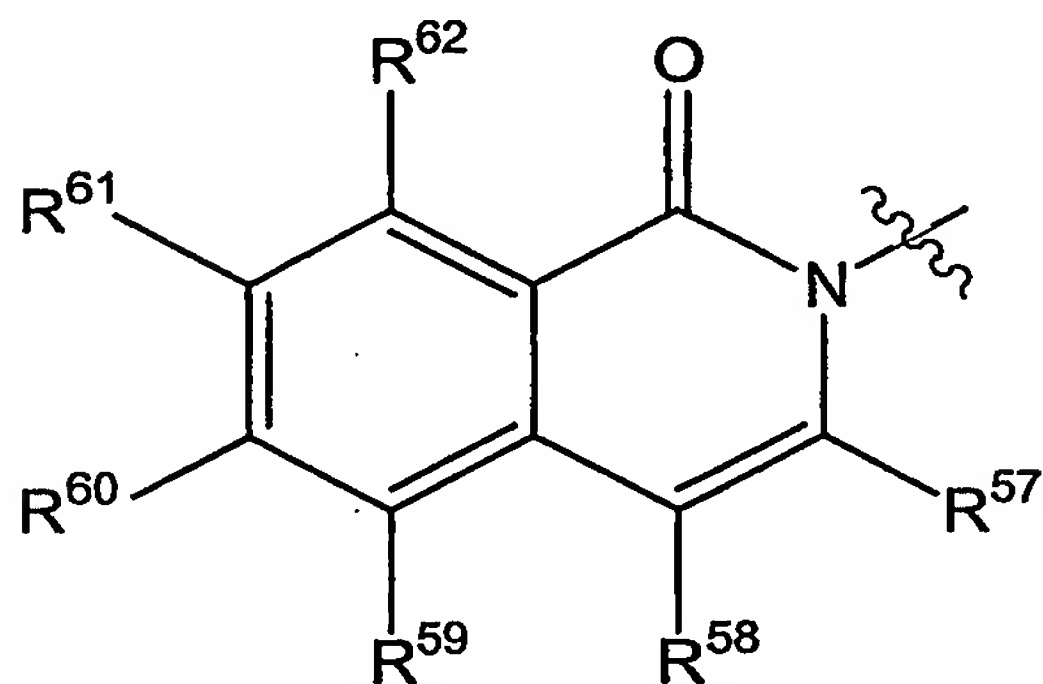


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12. The monomer of claim 1, wherein B is:

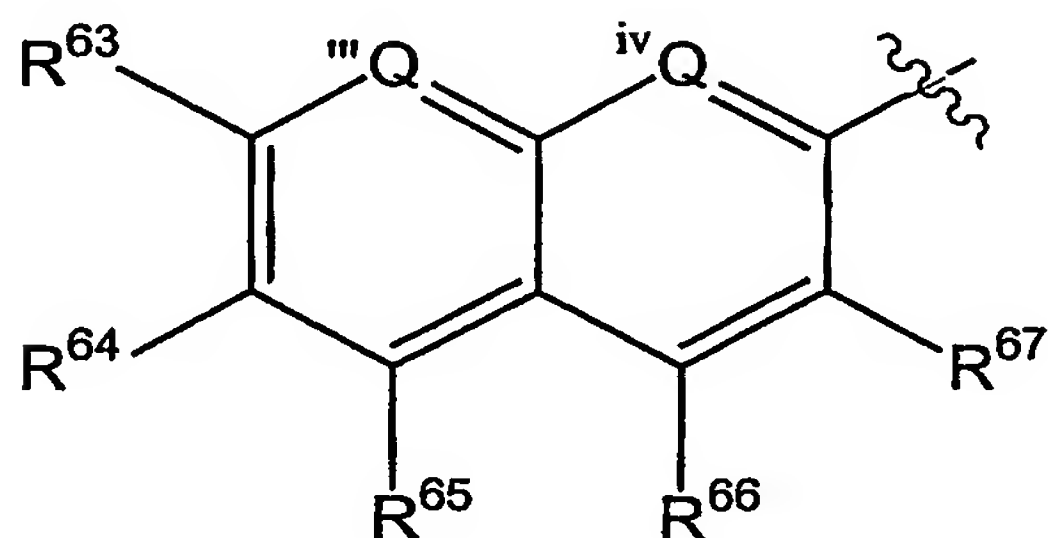


13. The monomer of claim 1, wherein B is:



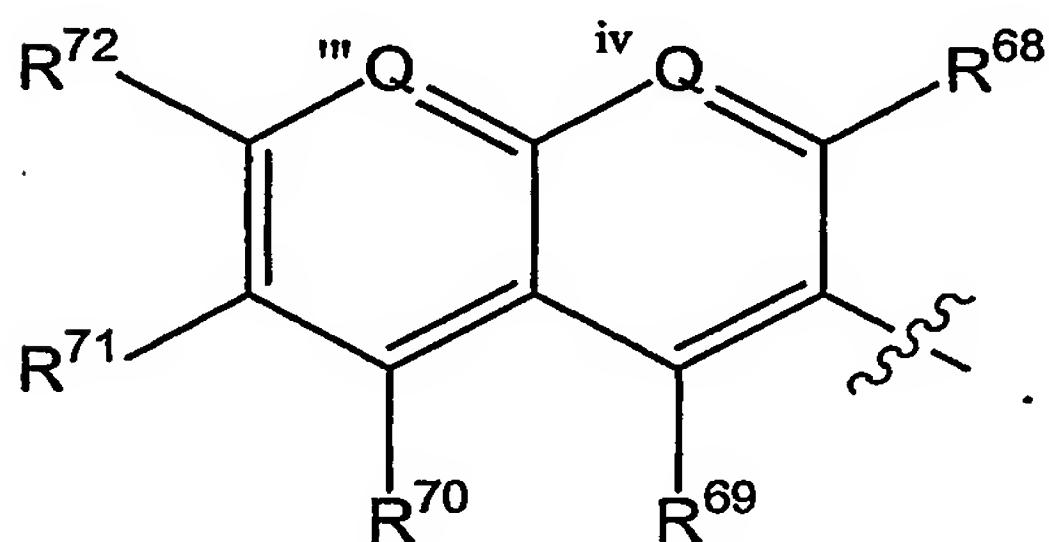
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14. The monomer of claim 1, wherein B is:



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15. The monomer of claim 1, wherein B is:



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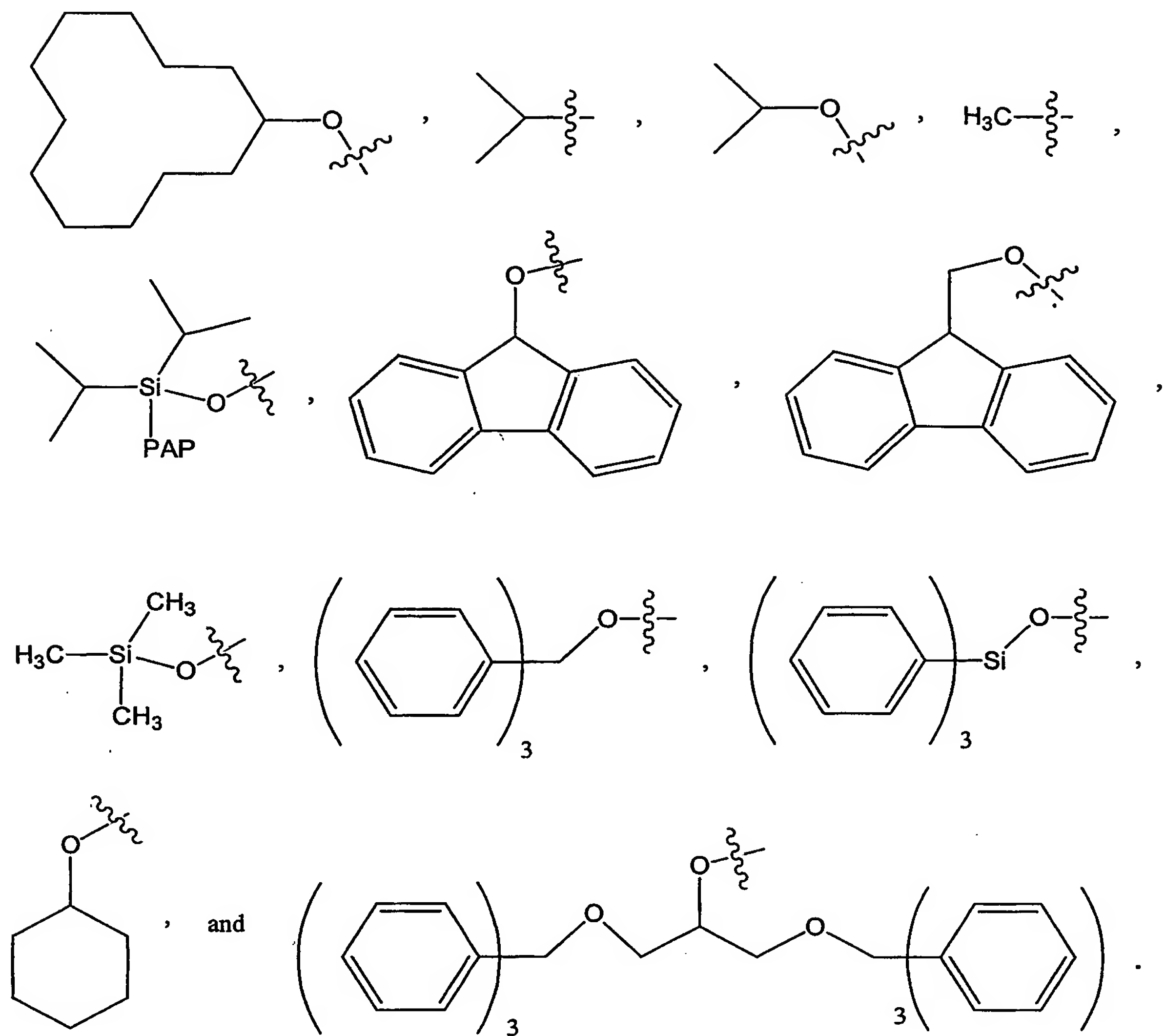
16. The monomer of claim 1, wherein B is anthracenyl.

17. The monomer of claim 1, wherein B is pyrenyl.

18. The monomer of claim 1, wherein R²⁸ is isopropyl.

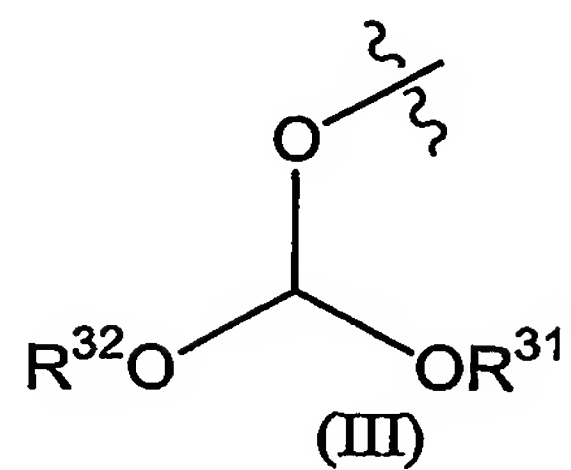
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19. The monomer of claim 1, wherein X^{5'}, X^{5''}, and X^{5'''} are any combination of the following formula:

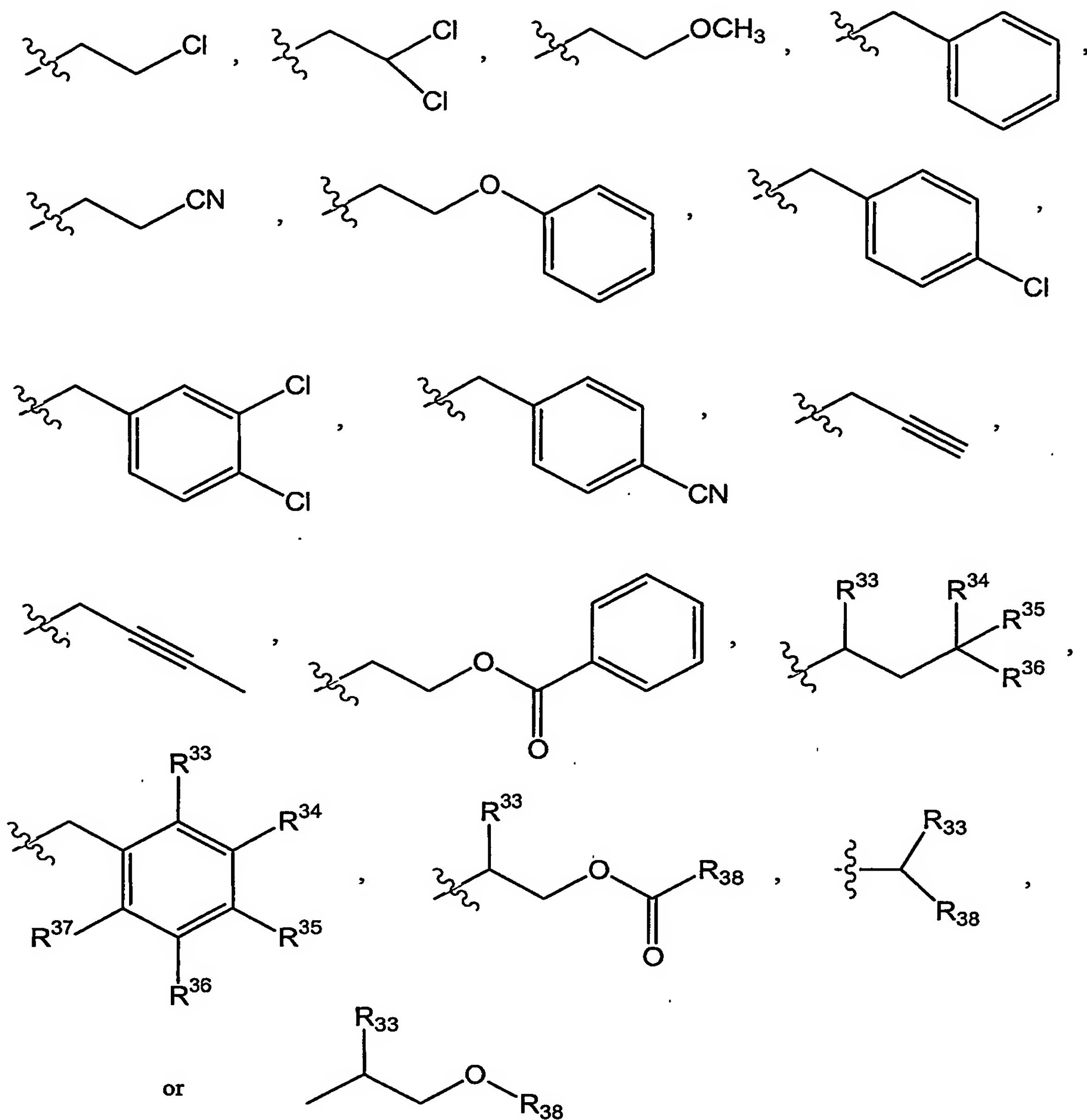


- 5 20. The compound of claim 1, wherein X^{5'} and X^{5''} are siloxy and X^{5'''} is cycloalkoxy.

21. The monomer of claim 1, wherein the orthoester protecting group has a formula(III):

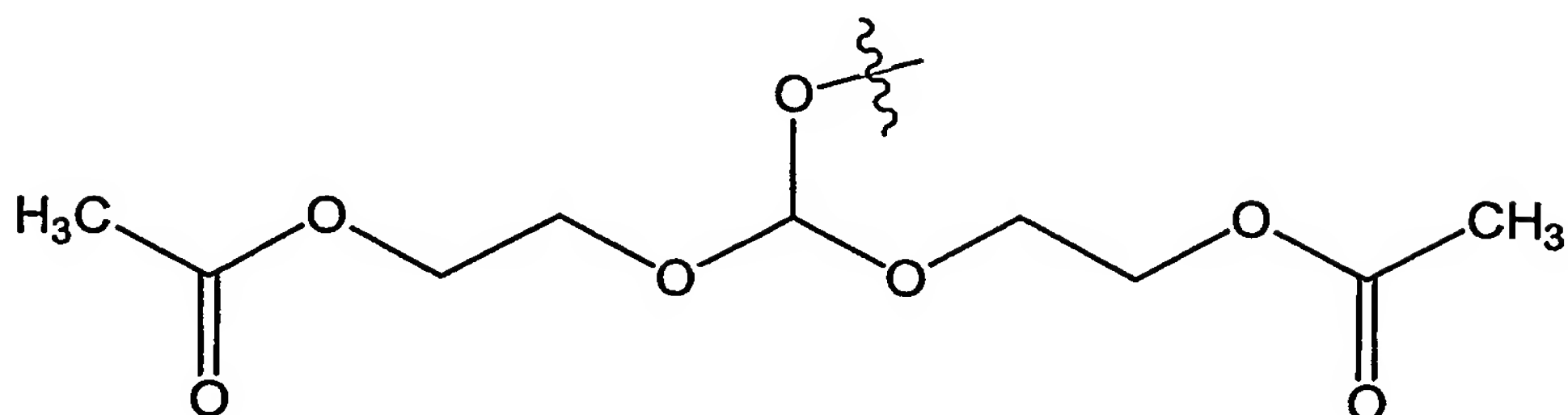


5 22. The monomer of claim 21, wherein R³¹ and R³² are the same or different and are any combination of the following formulae:



wherein R^{33} , R^{34} , R^{35} , R^{36} , and R^{37} is a compatible ligand, or hydrogen, or halogen, alkyl, or cyano substituent, and R^{38} is compatible ligand.

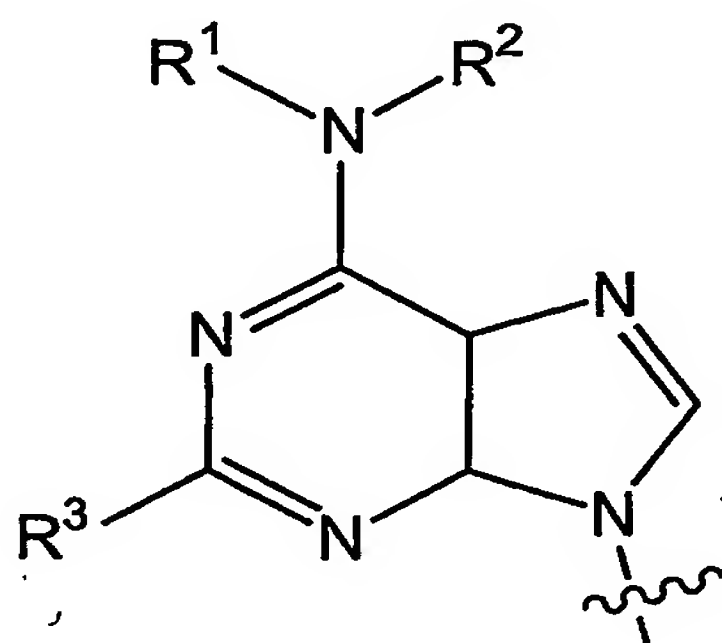
23. The monomer of claim 21, wherein the orthoester is:



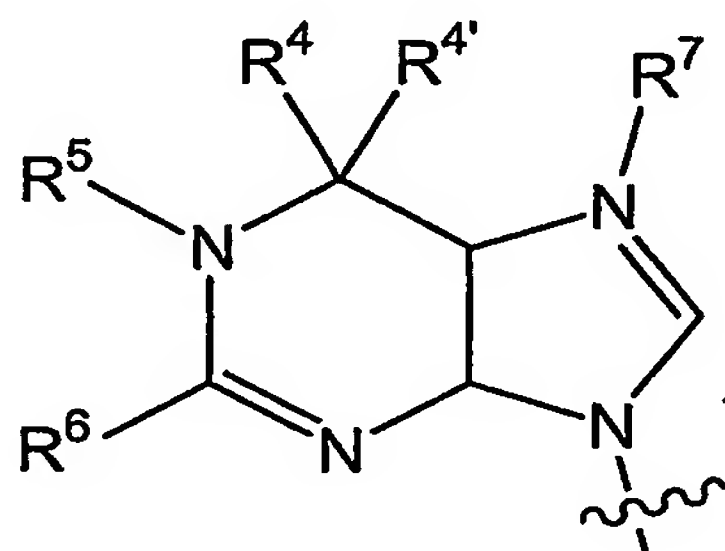
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24. The monomer of claim 1, wherein R^{29} is a fluoride-stable polystyrene based solid support or PEG.

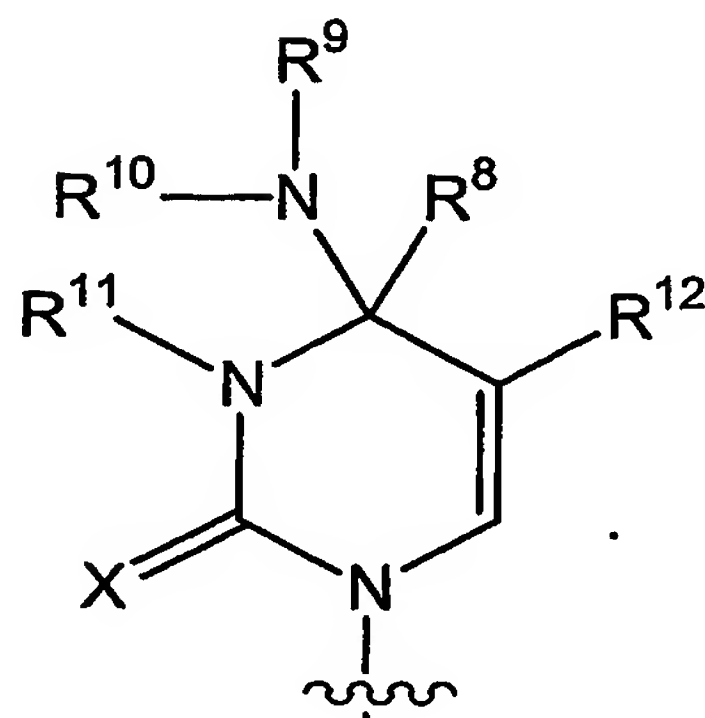
25. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ;
 10 R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



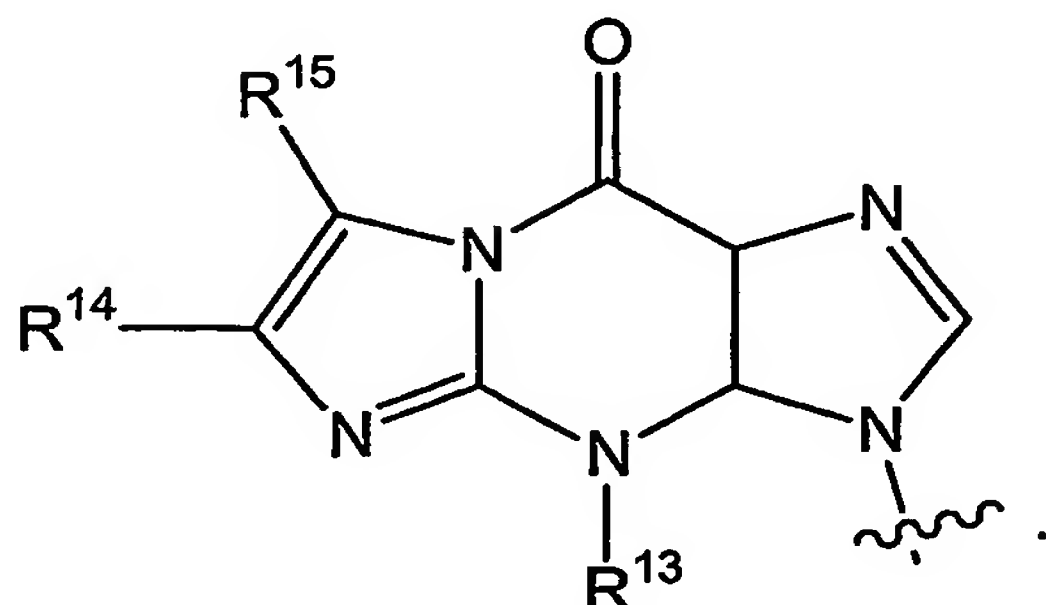
26. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ;
 15 R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



27. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is

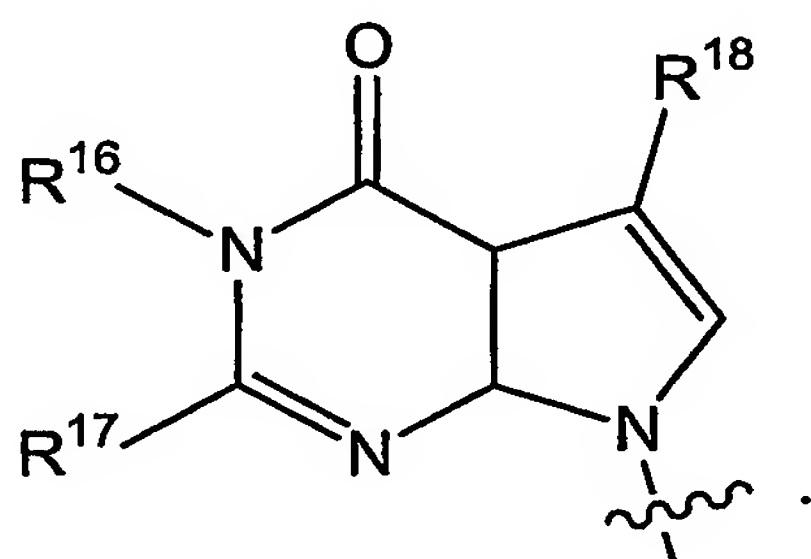


28. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



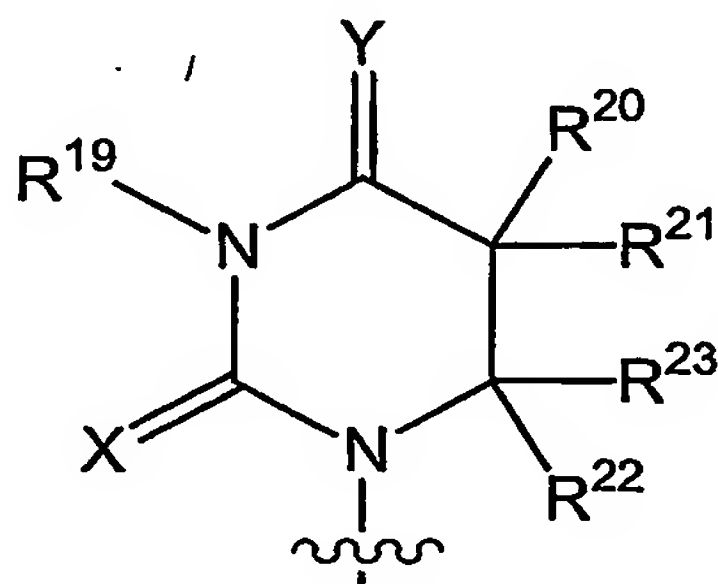
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29. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



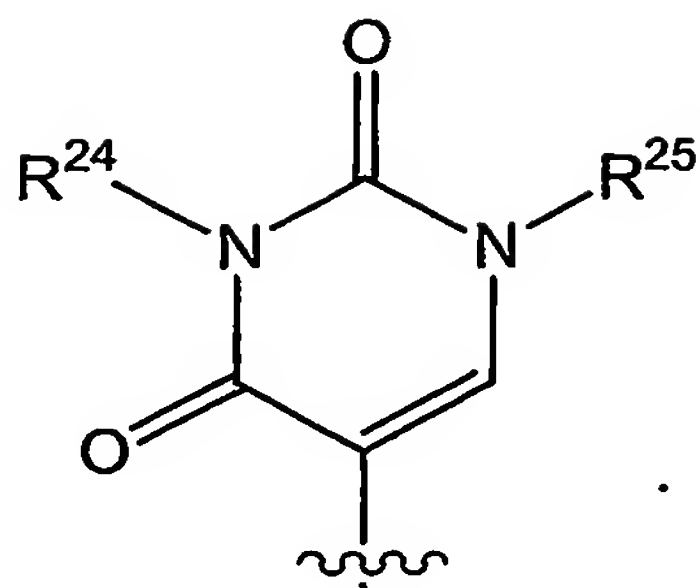
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30. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

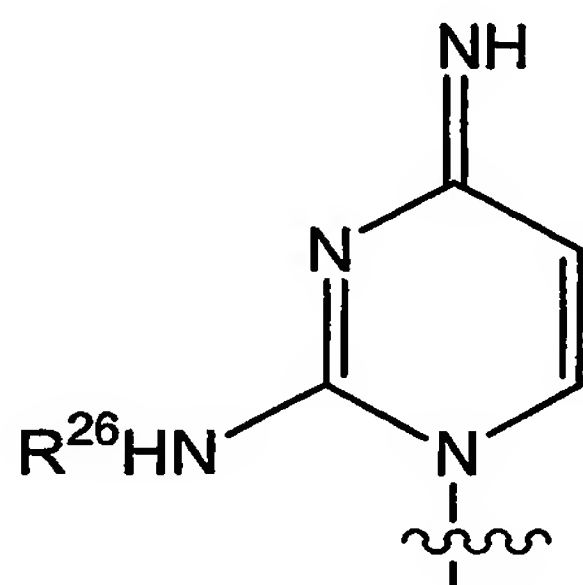


31. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

15

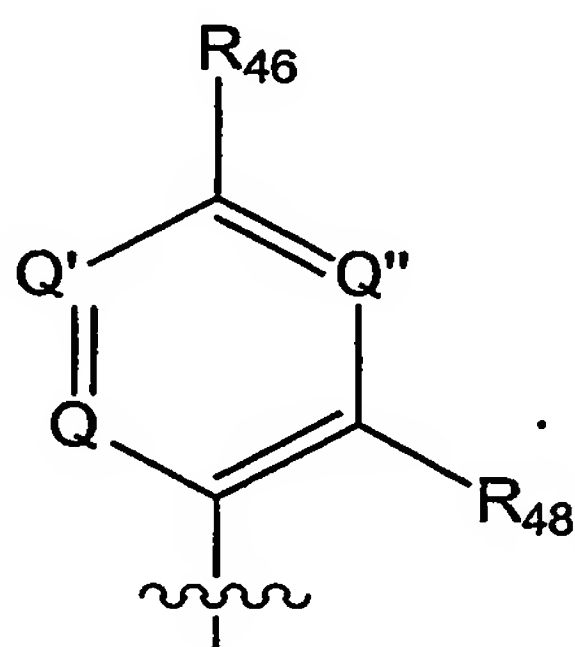


32. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



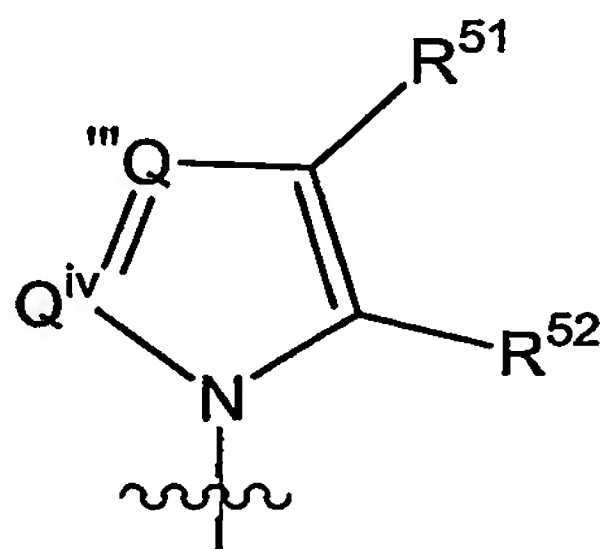
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33. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

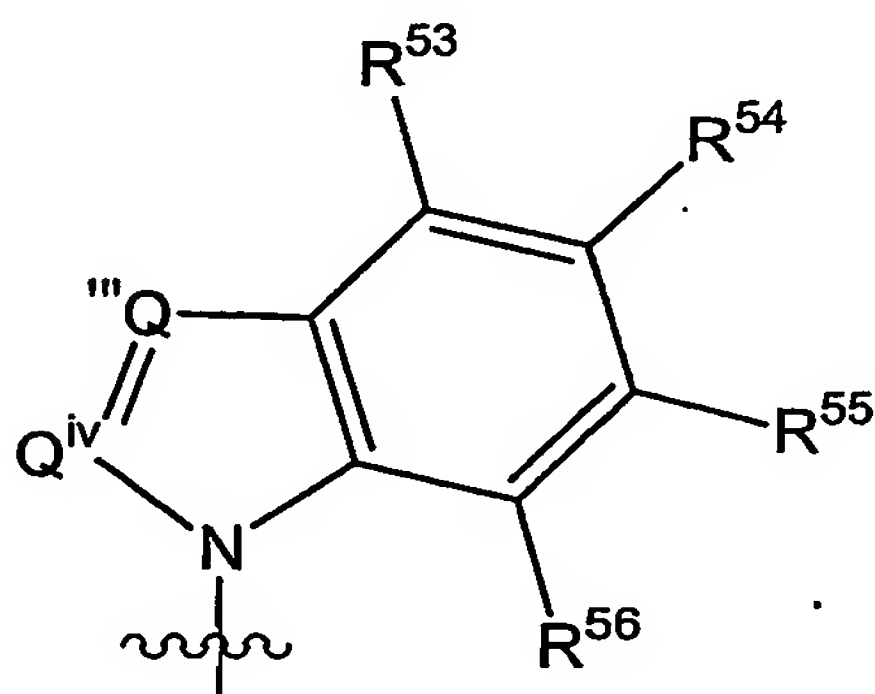


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34. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

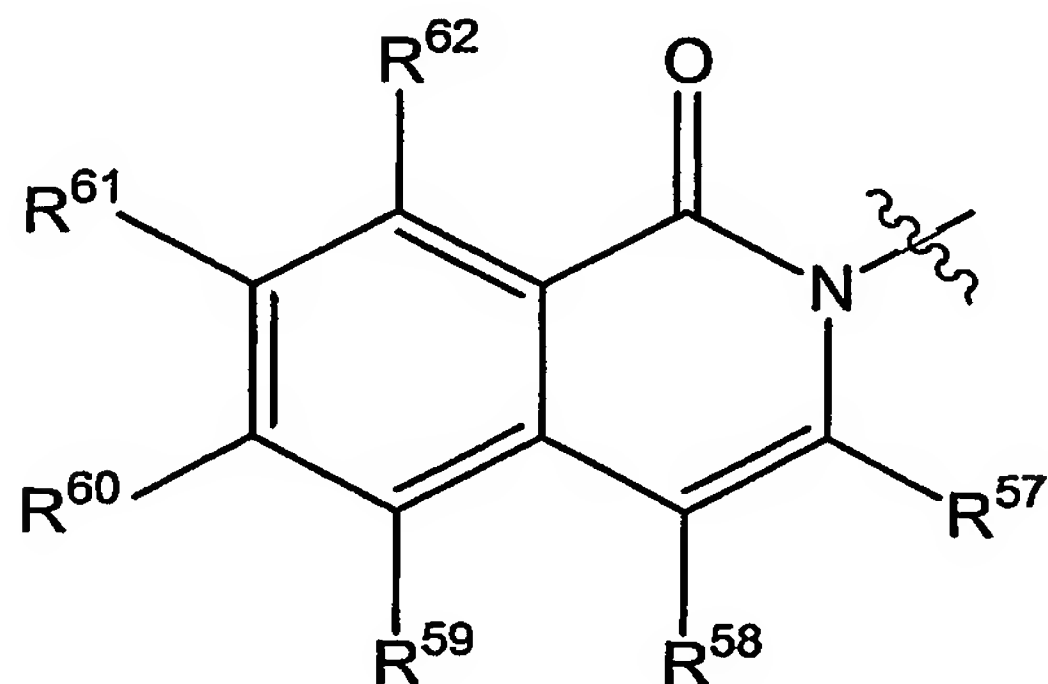


5 35. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

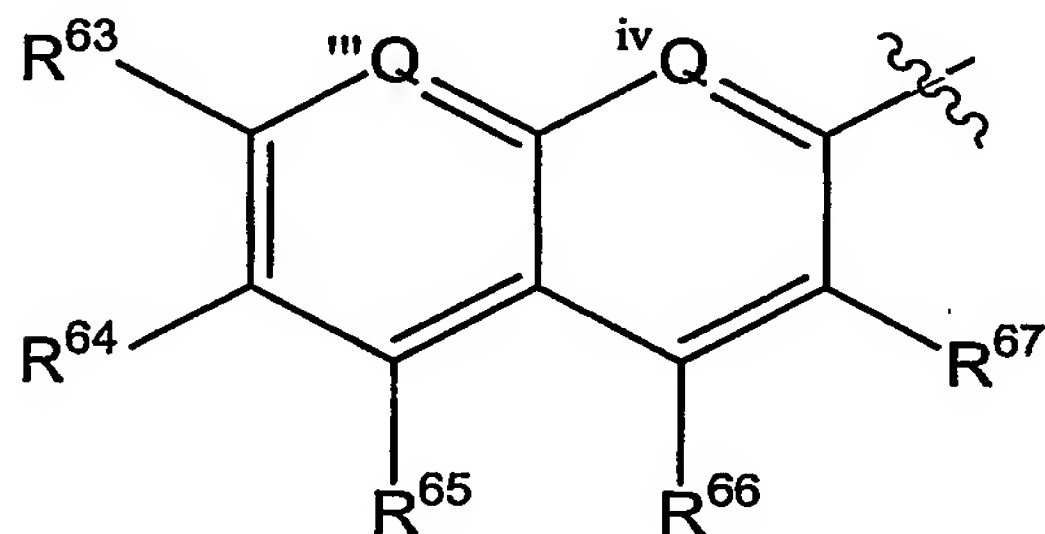


36. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:

5

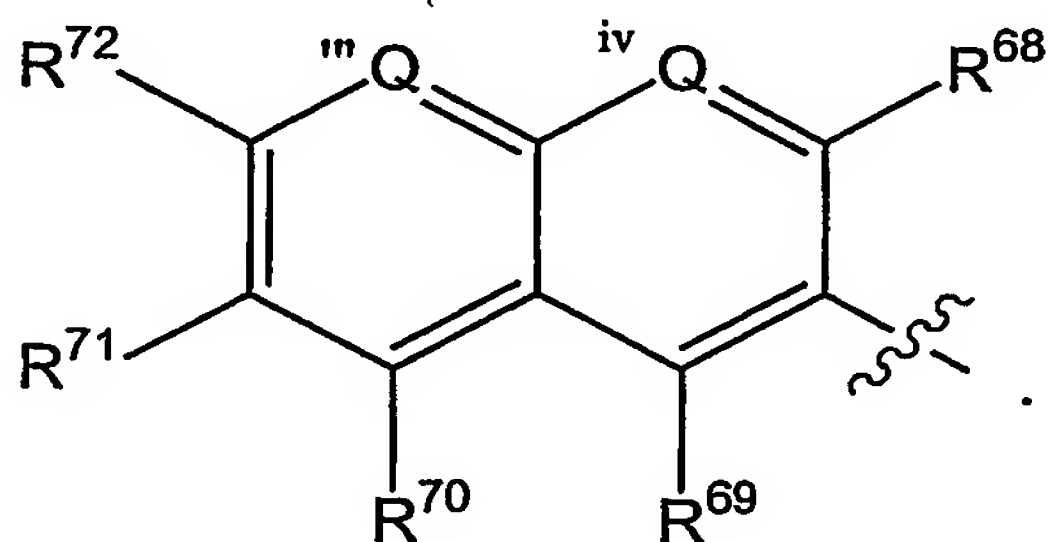


37. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



10

38. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is:



15

39. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is anthracenyl.

40. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is pyrenyl.

41. The monomer of claim 1, wherein B is selected from the group consisting of:

2-aminoadeninyl

2-methyladeninyl,

N6-methyladeninyl,

2-methylthio-N6-methyladeninyl,

N6-isopentenyladeninyl,

2-methylthio-N6-isopentenyladeninyl,

N6-(cis-hydroxyisopentenyl)adeninyl,

2-methylthio-N6-(cis-hydroxyisopentenyl) adeninyl,

N6-glycinylocarbamoyladeninyl,

N6-threonylocarbamoyladeninyl,

2-methylthio-N6-threonyl carbamoyladeninyl,

N6-methyl-N6-threonylocarbamoyladeninyl,

N6-hydroxynorvalylocarbamoyladeninyl,

2-methylthio-N6-hydroxynorvalyl carbamoyladeninyl,

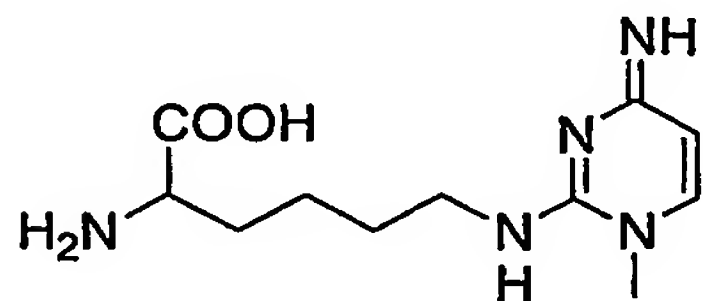
N6,N6-dimethyladeninyl,

3-methylcytosinyl,

5-methylcytosinyl,

2-thiocytosinyl,

5-formylcytosinyl,

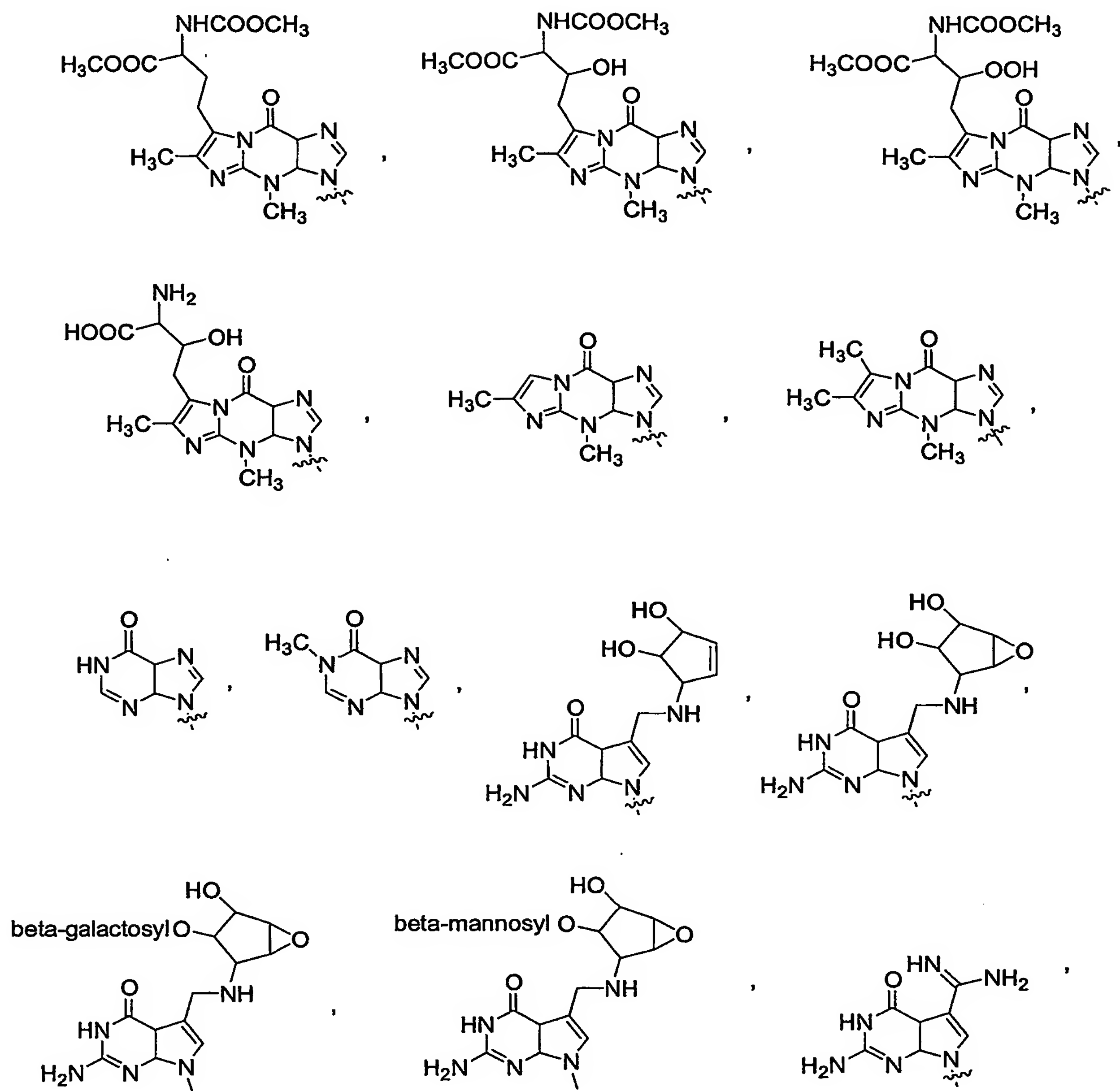


N4-methylcytosinyl,

5-hydroxymethylcytosinyl,

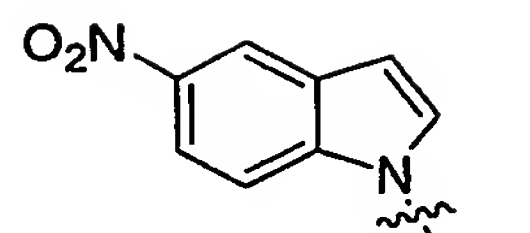
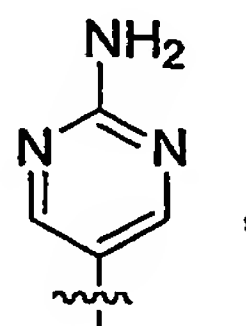
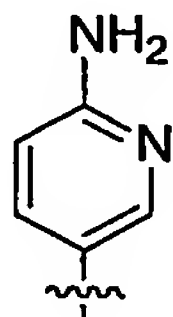
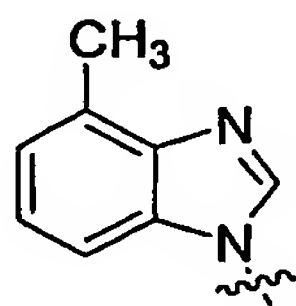
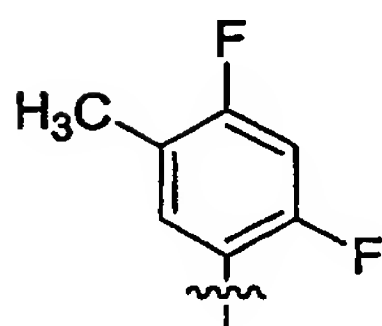
1-methylguaninyl,

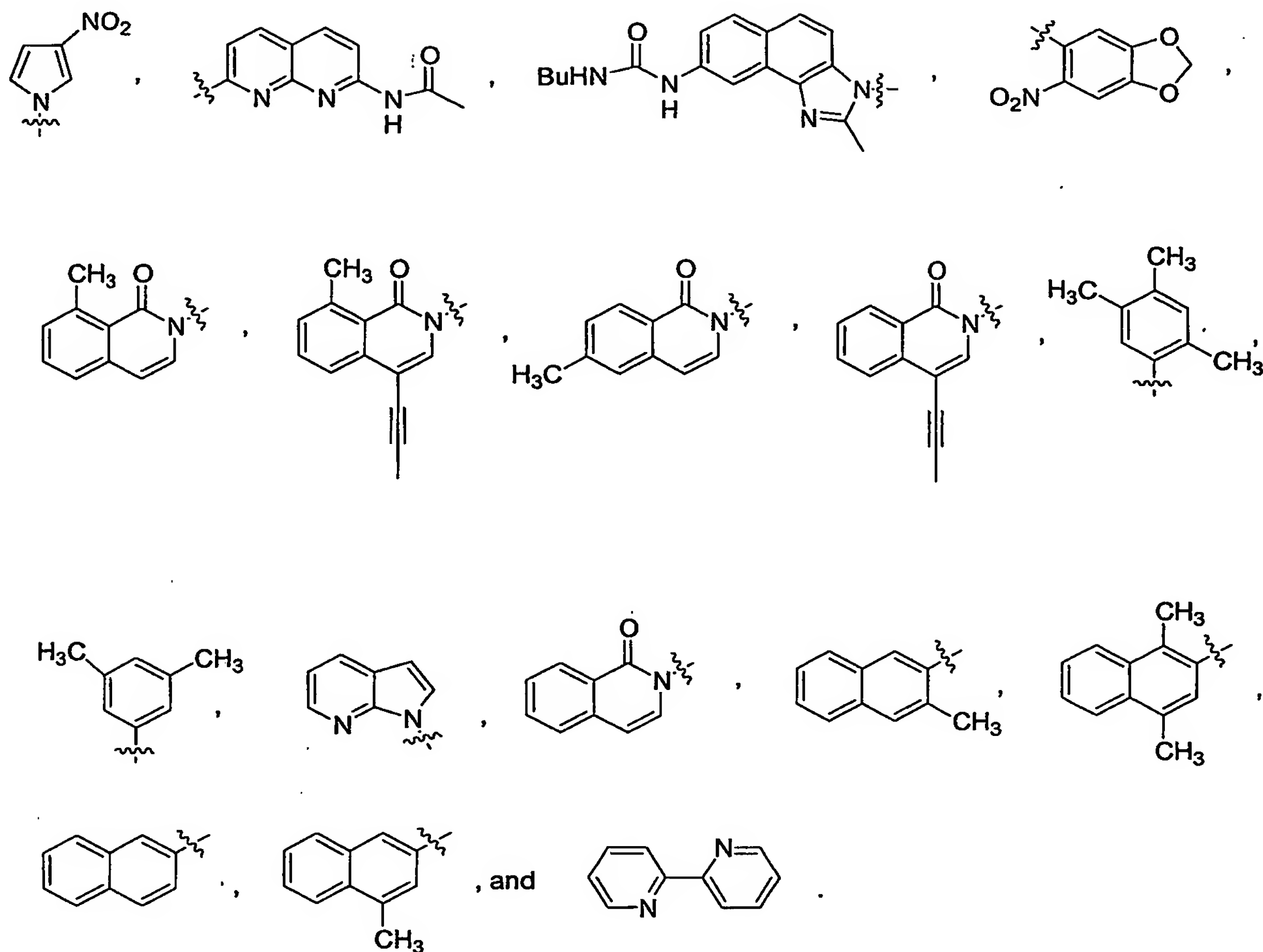
N2-methylguaninyl,
7-methylguaninyl,
N2,N2-dimethylguaninyl,



N2,7-dimethylguaninyl,
N2,N2,7-trimethylguaninyl,
1-methylguaninyl,
7-cyano-7-deazaguaninyl,
7-aminomethyl-7-deazaguaninyl,
pseudouracilyl,

- 5 dihydouracilyl,
5-methyluracilyl,
1-methylpseudouracilyl,
2-thiouracilyl,
4-thiouracilyl,
5-methyl-2-thiouracilyl,
3-(3-amino-3-carboxypropyl)uracilyl,
5-hydroxyuracilyl,
5-methoxyuracilyl,
10 uracilyl 5-oxyacetic acid,
uracilyl 5-oxyacetic acid methyl ester,
5-(carboxyhydroxymethyl)uracilyl,
5-(carboxyhydroxymethyl)uracilyl methyl ester,
5-methoxycarbonylmethyluracilyl,
15 5-methoxycarbonylmethyl-2-thiouracilyl,
5-aminomethyl-2-thiouracilyl,
5-methylaminomethyluracilyl,
5-methylaminomethyl-2-thiouracilyl,
5-methylaminomethyl-2-selenouracilyl,
20 5-carbamoylmethyluracilyl,
5-carboxymethylaminomethyluracilyl,
5-carboxymethylaminomethyl-2-thiouracilyl,
3-methyluracilyl,
1-methyl-3-(3-amino-3-carboxypropyl) pseudouracilyl,
25 5-carboxymethyluracilyl,
5-methyldihydouracilyl,
3-methylpseudouracilyl,





42. The monomer of claim 1, wherein X^2 is $-\text{OC}[\text{OCH}_2\text{CH}_2\text{OC}(\text{O})\text{CH}_3]_2$; R^{27} is CH_3 ; R^{28} is $(\text{CH}_3)_2\text{CH}-$; $X5'$ and $X5''$ are trimethylsiloxy; $X5'''$ is cyclododecyloxy; and B is selected from the group consisting of:

2-aminoadeninyl,

2-methyladeninyl,

N6-methyladeninyl,

2-methylthio-N6-methyladeninyl,

N6-isopentenyladeninyl,

2-methylthio-N6-isopentenyladeninyl,

N6-(cis-hydroxyisopentenyl)adeninyl,

2-methylthio-N6-(cis-hydroxyisopentenyl) adeninyl,

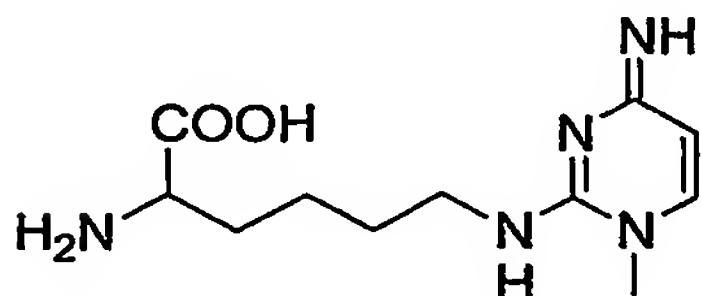
N6-glycinylocarbamoyladeninyl,

N6-threonylocarbamoyladeninyl,

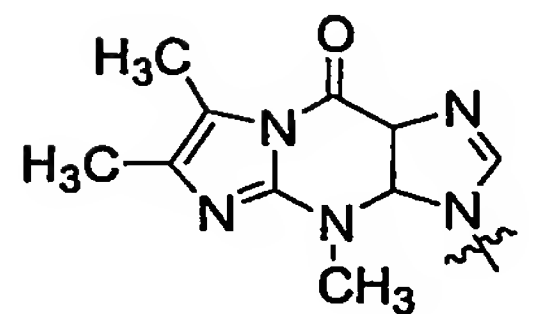
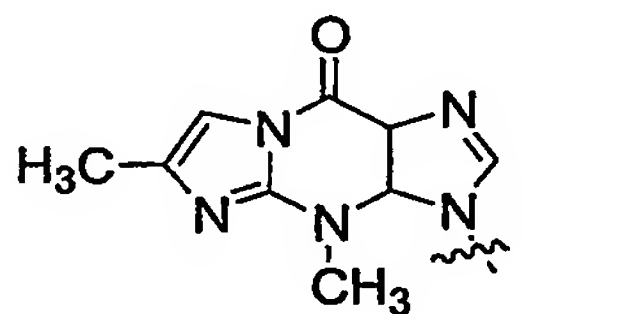
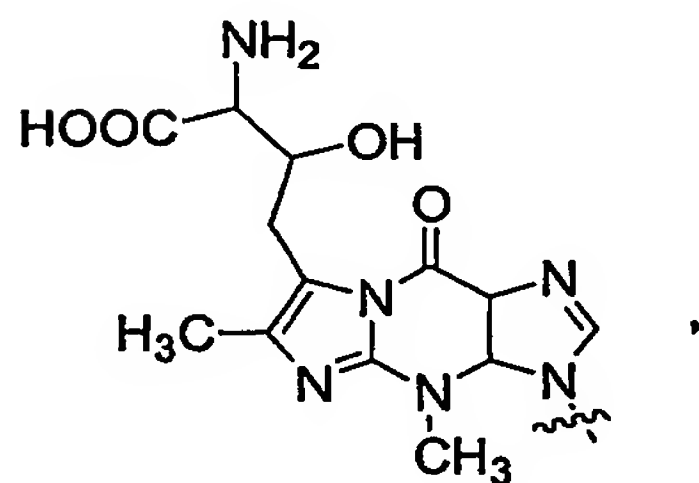
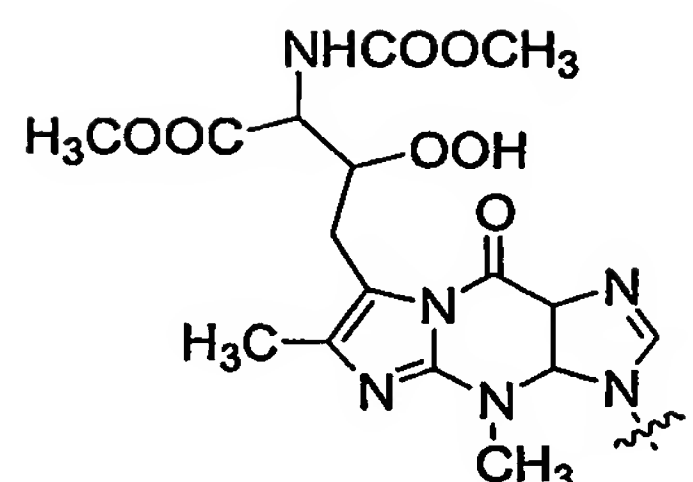
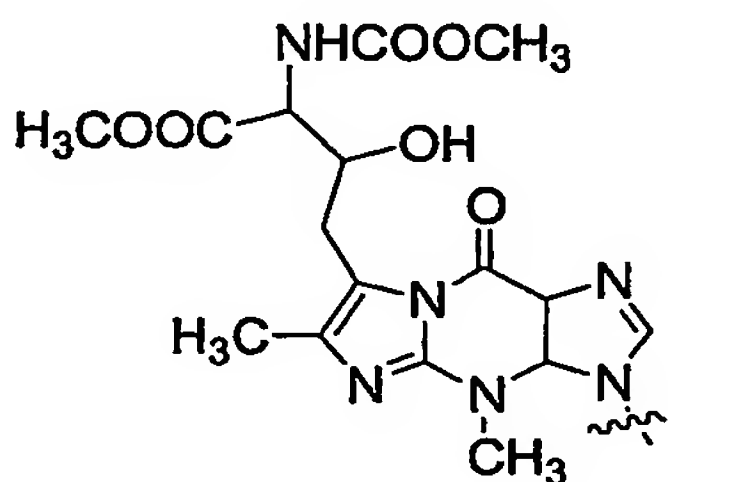
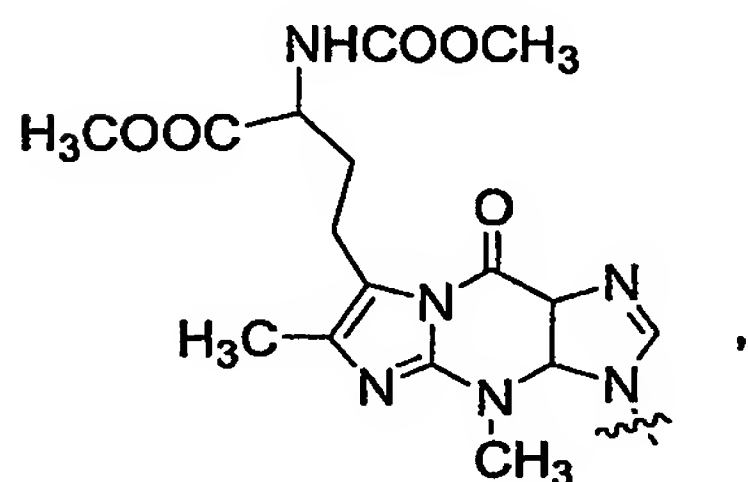
2-methylthio-N6-threonyl carbamoyladeninyl,

N6-methyl-N6-threonylocarbamoyladeninyl,

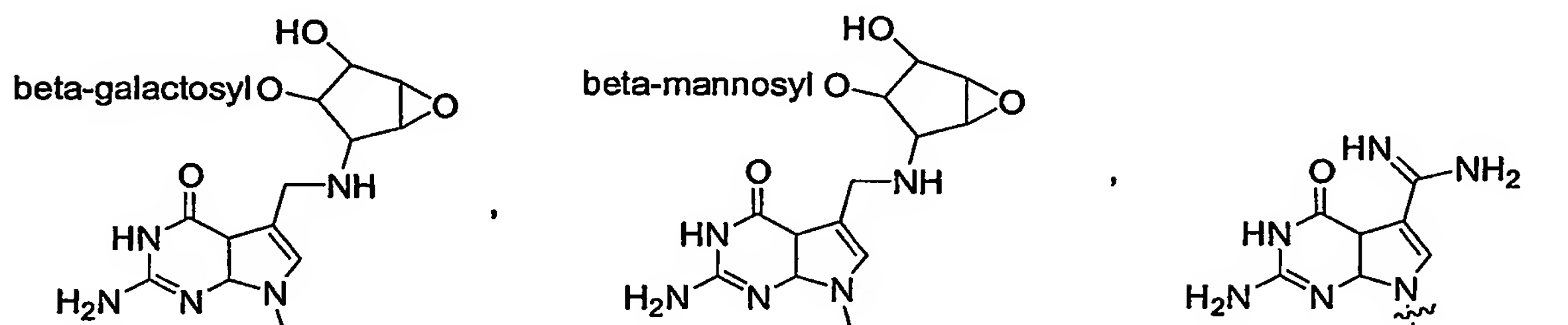
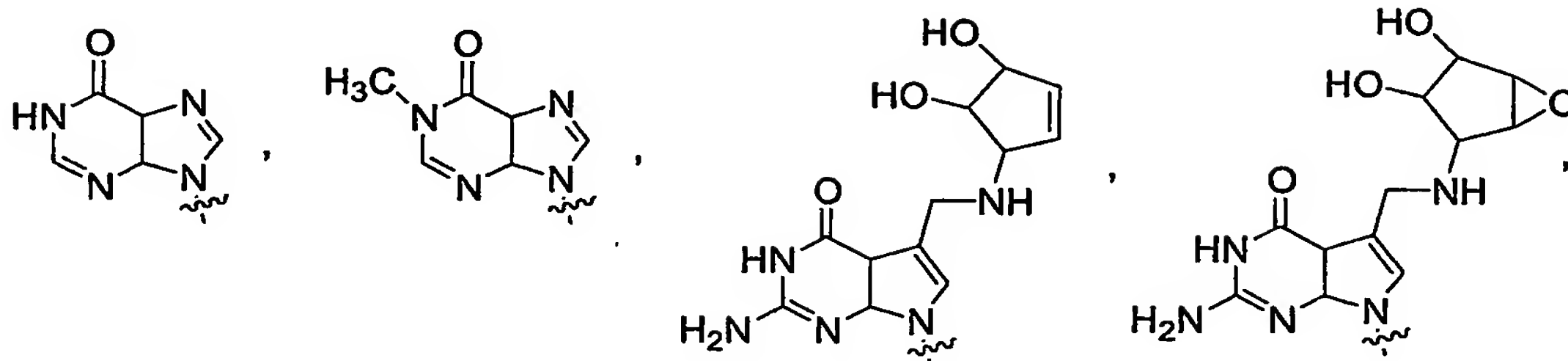
N6-hydroxynorvalylcarbamoyladeninyl,
 2-methylthio-N6-hydroxynorvalyl carbamoyladeninyl,
 N6,N6-dimethyladeninyl,
 3-methylcytosinyl,
 5-methylcytosinyl,
 2-thiocytosinyl,
 5-formylcytosinyl,



N4-methylcytosinyl,
 5-hydroxymethylcytosinyl,
 1-methylguaninyl,
 N2-methylguaninyl,
 7-methylguaninyl,
 N2,N2-dimethylguaninyl,

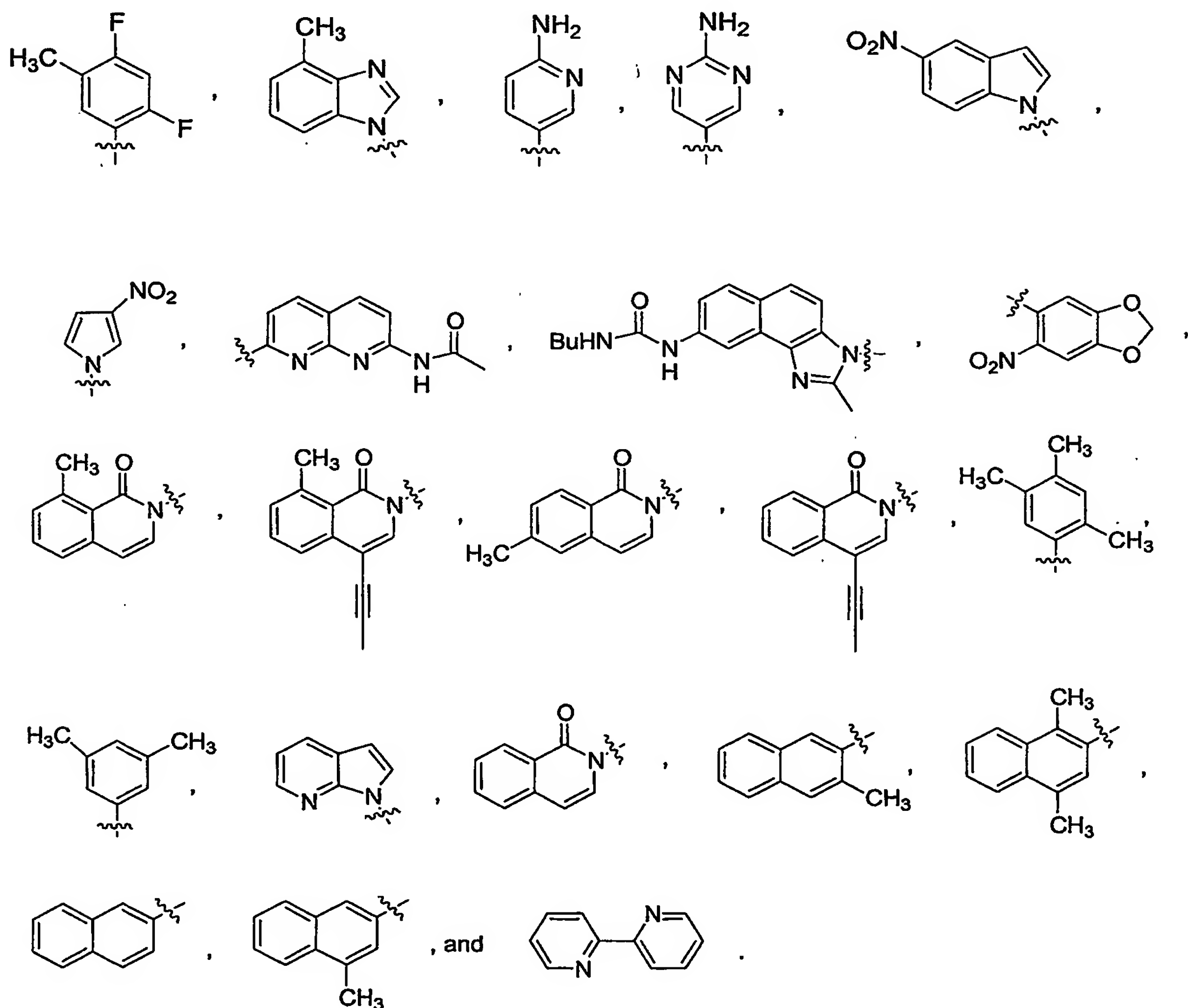


N2,7-dimethylguaninyl,
 N2,N2,7-trimethylguaninyl,
 1-methylguaninyl,
 5 7-cyano-7-deazaguaninyl,
 7-aminomethyl-7-deazaguaninyl,
 pseudouracilyl,
 dihydrouracilyl,
 5-methyluracilyl,
 10 1-methylpseudouracilyl,
 2-thiouracilyl,
 4-thiouracilyl



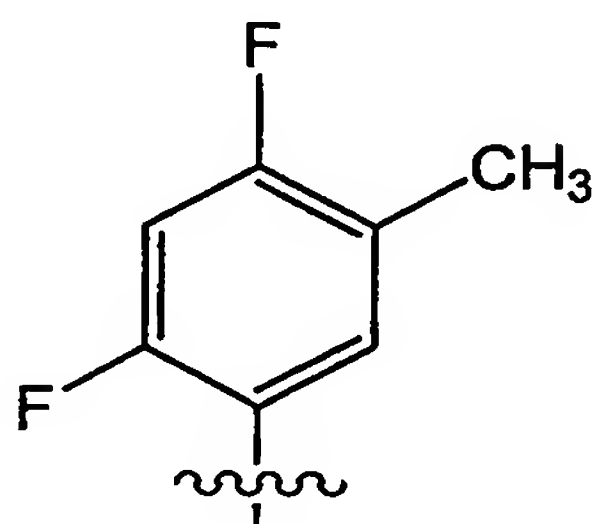
5-methyl-2-thiouracilyl,
 3-(3-amino-3-carboxypropyl)uracilyl,
 5-hydroxyuracilyl,
 5-methoxyuracilyl,
 uracilyl 5-oxyacetic acid,
 uracilyl 5-oxyacetic acid methyl ester,
 5-(carboxyhydroxymethyl)uracilyl,
 5-(carboxyhydroxymethyl)uracilyl methyl ester,
 5-methoxycarbonylmethyluracilyl,
 5-methoxycarbonylmethyl-2-thiouracilyl,

5-aminomethyl-2-thiouracilyl,
5-methylaminomethyluracilyl,
5-methylaminomethyl-2-thiouracilyl,
5-methylaminomethyl-2-selenouracilyl,
5
5-carbamoylmethyluracilyl,
5-carboxymethylaminomethyluracilyl,
5-carboxymethylaminomethyl-2-thiouracilyl,
3-methyluracilyl,
1-methyl-3-(3-amino-3-carboxypropyl) pseudouracilyl,
10
5-carboxymethyluracilyl,
5-methyldihydrouracilyl,
3-methylpseudouracilyl,



43. The monomer of claim 1, wherein X^2 is fluoro.

44. The monomer of claim 1, wherein B is:

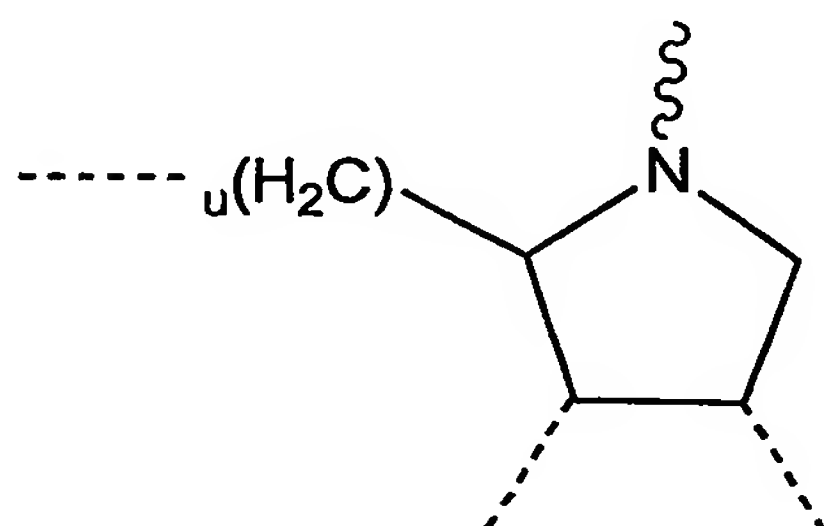


5

45. The monomer of claim 1, wherein B is substituted or unsubstituted aryl attached to a tethered or untethered ligand.

10

46. A protected monomer having a formula:

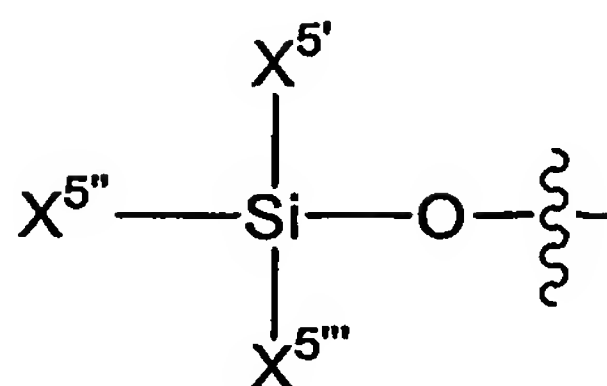


in which

u is 1 or 2; the wavy line represents a point of attachment for a ligand or a tethered ligand; and the dotted lines represent points of attachment for a first functionalized hydroxyl group; a second functionalized hydroxyl group; and an unfunctionalized hydroxyl group, a protected hydroxyl group, or hydrogen.

15

47. The monomer of claim 46, wherein the first functionalized hydroxyl group has the formula:

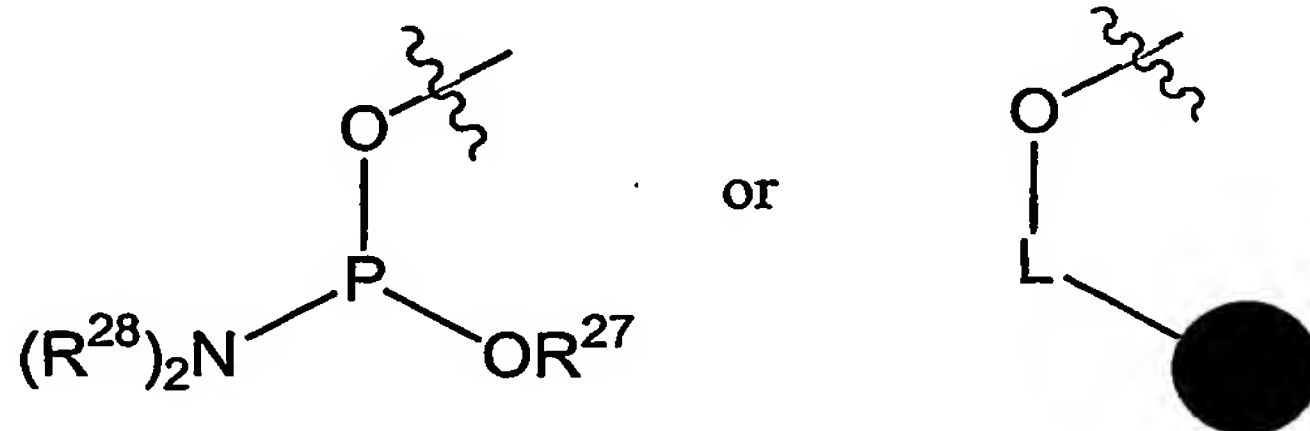


5

; in which

$X^{5'}$, $X^{5''}$, and $X^{5'''}$ include at least one alkoxy or siloxy substituent.

48. The monomer of claim 46, wherein the second functionalized hydroxyl group has one of the following formulas:



10

; in which

R^{27} is $\text{C}_1\text{-C}_6$ alkyl optionally substituted with cyano or $\text{C}_2\text{-C}_6$ alkenyl; R^{28} is $\text{C}_1\text{-C}_{10}$ alkyl; \bullet is a solid or liquid support reagent; and L is a linker.

15

49. The monomer of claim 46, wherein the ligand is a targeting group.

50. The monomer of claim 49, wherein the targeting group is a lipid, steroid, vitamin, carbohydrate, polyamine, amino acid, peptide, peptide mimetic or cleaving molecule.

20

51. The monomer of claim 50, wherein the steroid is cholesterol.

52. The monomer of claim 46, wherein the ligand is a diagnostic group.

5 53. The monomer of claim 52, wherein the diagnostic group is biotin, a fluorophore, an antibody or an antigen.

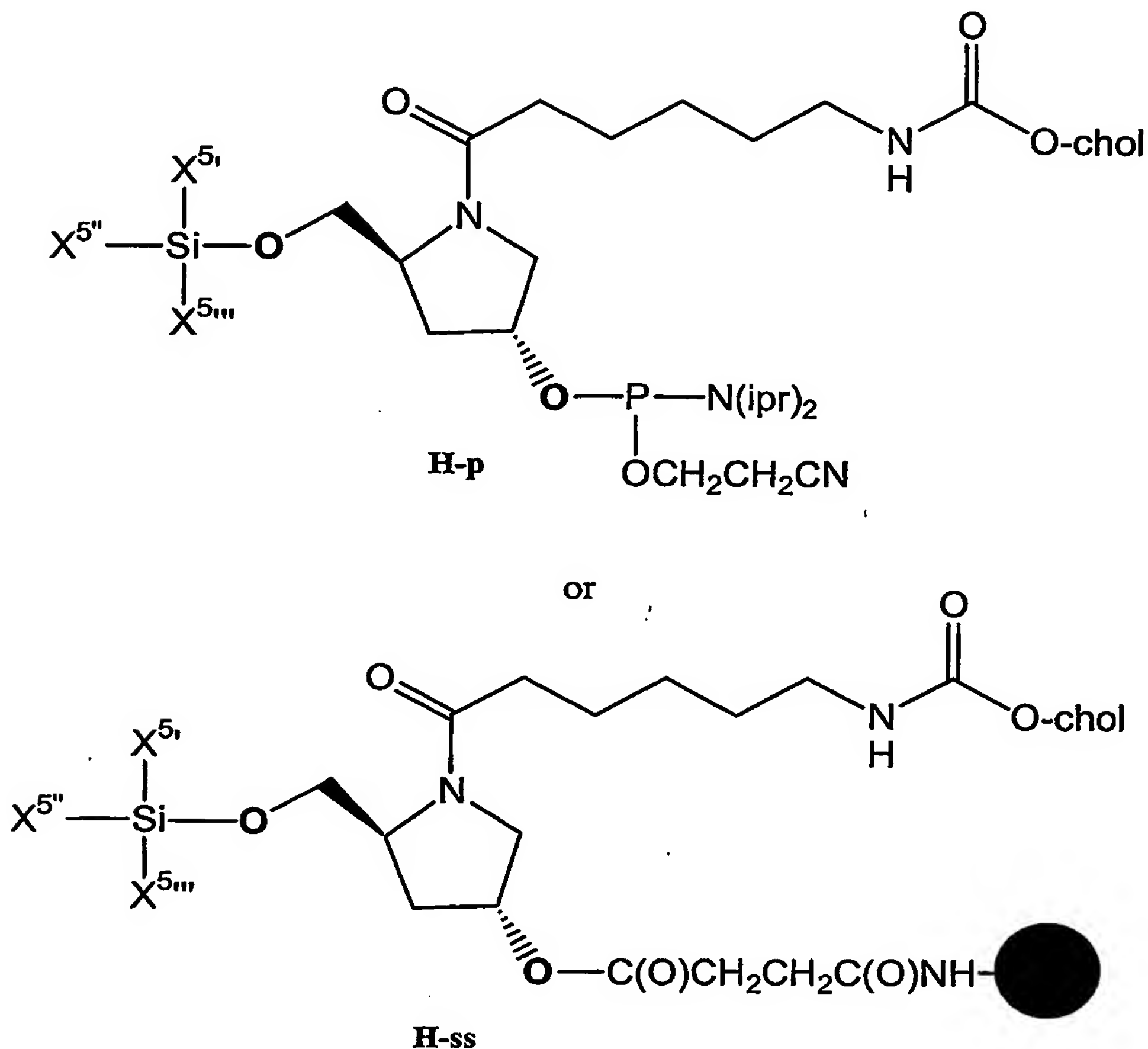
10 54. The monomer of claim 46, wherein the ligand has a formula $(G)C(=H)NHR^n$, in which G is -O-, -NH-, or -CH₂-; H is O or NH; and Rⁿ is H, C₁-C₆ alkyl, C₆-C₁₀ aryl, or C₅-C₁₀ heteroaryl.

55. The monomer of claim 46, wherein the monomer has a tethered ligand.

15 56. The monomer of claim 55, wherein the ligand is tethered with a tether selected from the group consisting of: -C(O)-(CH₂)_s-C(O)-(ligand); -C(O)-(CH₂)_s-C(O)O-(ligand); -C(O)-O-(ligand); -C(O)-(CH₂)_s-NH-; -C(O)-(CH₂)_s-NH-C(O)-(ligand); -C(O)-(CH₂)_s-(ligand); -C(O)-NH-(ligand); -C(O)-(ligand); -(CH₂)_s-C(O)-(ligand); -(CH₂)_s-C(O)O-(ligand); -(CH₂)_s-(ligand); -(CH₂)_s-NH-; and -(CH₂)_s-NH-C(O)-(ligand), wherein s is 0-6.

20

57. The monomer of claim 46, wherein the monomer has the formula:



5 wherein, $\text{X}^{5'}$, $\text{X}^{5''}$, and $\text{X}^{5'''}$ include at least one alkoxy or siloxy substituent, ipr is an isopropyl group, and chol is a cholesterol radical.

58. An iRNA agent having a monomer of claim 1 or 46.

10 59. A method of making an iRNA agent, the method comprising providing an iRNA agent having a monomer of claim 1 or 46 and allowing it to anneal to a complementary RNA sequence to form an iRNA agent.